

**WEST**[Help](#)[Logout](#)[Interrupt](#)[Main Menu](#)[Search Form](#)[Posting Counts](#)[Show S Numbers](#)[Edit S Numbers](#)[Preferences](#)**Search Results -**

Terms	Documents
(crustacean? and fish)and (feedstuff)	19

Database: US Patents Full-Text Database  
US Pre-Grant Publication Full-Text Database  
JPO Abstracts Database  
EPO Abstracts Database  
Derwent World Patents Index  
IBM Technical Disclosure Bulletins

Refine Search: (crustacean? and fish)and (feedstuff) Clear

**Search History****Today's Date: 4/3/2001**

<u>DB Name</u>	<u>Query</u>	<u>Hit Count</u>	<u>Set Name</u>
USPT	(crustacean? and fish)and (feedstuff)	19	<u>L13</u>
USPT	(crustaceans? and fish)and (feedstuff)	0	<u>L12</u>
USPT	(crustaceans? and fish)and (feed)	0	<u>L11</u>
USPT	(crustaceans?)and (immunity)	0	<u>L10</u>
USPT	(crustaceans?)and (pantoea)	0	<u>L9</u>
USPT	(crustaceans?)and (gram negative bacteria?)	2	<u>L8</u>
USPT	(crustacens?)and (gram negative bacteria?)	0	<u>L7</u>
USPT	(crustacens?)and (pantoea)	0	<u>L6</u>
USPT	(crustac?)and (pantoea)	0	<u>L5</u>
USPT	(gram negative bacteria) AND (fish)	16194	<u>L4</u>
USPT	(gram negative bacteria) AND (fish)	16194	<u>L3</u>
USPT	(gram negative bacteria)[AB] AND (fish)[AB]	65	<u>L2</u>
USPT	(gram negative bacteria)[AB] AND (fish)	945	<u>L1</u>

**WEST****Freeform Search****Database:**

US Patents Full-Text Database  
US Pre-Grant Publication Full-Text Database  
JPO Abstracts Database  
EPO Abstracts Database  
Derwent World Patents Index  
IBM Technical Disclosure Bulletins

**Term:**

(crustacean? and fish)and (lipopolysaccharide)

**Display:****Documents in Display Format:****Starting with Number****Generate:** ☐ Hit List ☒ Hit Count ☐ Image**Search History****Today's Date:** 4/3/2001

<u>DB Name</u>	<u>Query</u>	<u>Hit Count</u>	<u>Set Name</u>
USPT	(crustacean? and fish)and (lipopolysaccharide)	8	<u>L14</u>
USPT	(crustacean? and fish)and (feedstuff)	19	<u>L13</u>
USPT	(crustaceans? and fish)and (feedstuff)	0	<u>L12</u>
USPT	(crustaceans? and fish)and (feed)	0	<u>L11</u>
USPT	(crustaceans?)and (immunity)	0	<u>L10</u>
USPT	(crustaceans?)and (pantoea)	0	<u>L9</u>
USPT	(crustaceans?)and (gram negative bacteria?)	2	<u>L8</u>
USPT	(crustacens?)and (gram negative bacteria?)	0	<u>L7</u>
USPT	(crustacens?)and (pantoea)	0	<u>L6</u>
USPT	(crustac?)and (pantoea)	0	<u>L5</u>
USPT	(gram negative bacteria) AND (fish)	16194	<u>L4</u>
USPT	(gram negative bacteria) AND (fish)	16194	<u>L3</u>
USPT	(gram negative bacteria)[AB] AND (fish)[AB]	65	<u>L2</u>
USPT	(gram negative bacteria)[AB] AND (fish)	945	<u>L1</u>

[Generate Collection](#)[Print](#)**Search Results - Record(s) 1 through 10 of 42 returned.**

- 
- ☐ 1. [5494819](#). 12 Apr 94; 27 Feb 96. Pure culture of *Pantoea agglomerans* ferm BP-3511. [Soma](#); Gen-Ichiro, et al. 435/252.1; 435/243 435/253.6 514/54 514/885. C12N001/00 C12N001/20.
- 
- ☐ 2. [5413993](#). 20 Aug 92; 09 May 95. Method for the treatment of narcotic withdrawal symptoms in animals using lipopolysaccharides. [Soma](#); Gen-Ichiro, et al. 514/54; 424/93.4 424/93.48 435/252.31 435/252.5 435/822 514/885. A61K031/715 A61K037/00 C12N001/20.
- 
- ☐ 3. [5382430](#). 14 Dec 92; 17 Jan 95. Method for the stimulation of the immune system of an animal by use of limulus test-positive plant glycolipid. [Soma](#); Gen-Ichiro, et al. 424/195.15; 424/195.17 424/750 514/783. A61K035/78.
- 
- ☐ 4. [5346891](#). 20 Aug 91; 13 Sep 94. Lipopolysaccharide-producing bacteria, lipopolysaccharides, and lipopolysaccharide-containing, medicines and veterinary medicines. [Soma](#); Gen-Ichiro, et al. 514/54; 435/243 435/252.1 435/252.5 514/885. A61K031/715 C12N001/00 C12N001/20.
- 
- ☐ 5. [5281583](#). 22 Aug 91; 25 Jan 94. LPS-containing analgesics and veterinary analgesics. [Soma](#); Gen-Ichiro, et al. 514/54;. A61K031/70 A61K031/715.
- 
- ☐ 6. [5236709](#). 06 Feb 90; 17 Aug 93. Limulus test-positive plant glycolipid and method of stimulating the immunity system of an animal. [Soma](#); Gen-Ichiro, et al. 424/195.15; 424/195.17 424/750 514/783. A61R035/78.
- 
- ☐ 7. [5081021](#). 04 Feb 87; 14 Jan 92. DNA encoding HTNF variants exhibiting enhanced activity. [Mizuno](#); Den'ichi, et al. 435/69.5; 435/320.1 435/91.41 536/23.1 536/23.5. C12P021/00 C12P021/02 C07H015/12.
- 
- ☐ 8. [JP 08245702 A](#). 13 Mar 95. 24 Sep 96. HIGH-MOLECULAR WEIGHT LIPOPOLYSACCHARIDE. [MIZUNO](#), [DENICHI](#), et al. C08B037/00; C12P019/04.
- 
- ☐ 9. [JP 07109285 A](#). 24 May 91. 25 Apr 95. LIMULUS TEST-POSITIVE PLANT LPS, IMMUNOACTIVATOR, ANTIDIABETIC, ANALGESIC, AND THOSE FOR ANIMALS. [SOMA](#), [GENICHIRO](#), et al. C07G017/00; A61K035/78 A61K035/78 A61K035/78 A61K031/715 C08B037/00.
- 
- ☐ 10. [JP 06263650 A](#). 02 Dec 91. 20 Sep 94. LPS-CONTAINING ANTITUMOR AGENT AND ANTITUMOR AGENT FOR ANIMAL. [SOMA](#), [GENICHIRO](#), et al. A61K037/20; A61K035/64 A61K035/72 A61K035/78 A61K035/80 A61K035/82.
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Terms	Documents
L3 and LPS	42

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- 
- ☐ 11. JP 06141849 A. 30 Oct 92. 24 May 94. LPS PRODUCING BACTERIUM, LPS, IMMUNOLOGICAL FUNCTION ACTIVATOR AND IMMUNOLOGICAL FUNCTION ACTIVATOR FOR ANIMAL. SOMA, GENICHIRO, et al. C12N001/20; A23K001/16 A61K037/20 C08B037/00.
- 
- ☐ 12. JP 06090745 A. 19 Nov 92. 05 Apr 94. LPS-PRODUCING MICROORGANISM, LPS AND PHARMACEUTICAL AND ANIMAL DRUG CONTAINING LPS. SOMA, GENICHIRO, et al. 424/282.1 435/72. C12N001/20; A61K035/74 A61K037/20 A61K037/20 A61K037/20 A61K037/20 C08B037/00 C12P019/04.
- 
- ☐ 13. JP 06078756 A. 20 Aug 91. 22 Mar 94. MICROORGANISM CAPABLE OF PRODUCING LPS, LPS AND MEDICINE AND ANIMAL MEDICINE CONTAINING LPS. SOMA, GENICHIRO, et al. 424/278.1 435/252.1. C12N001/20; A61K037/20 A61K037/20 A61K037/20 A61K037/20 C08B037/00 C12P019/04.
- 
- ☐ 14. JP 06065092 A. 20 Aug 92. 08 Mar 94. ANTI-WITHDRAWAL SYMPTOMATIC AGENT CONTAINING LPS AND ANTI-WITHDRAWAL SYMPTOMATIC AGENT FOR ANIMAL. SOMA, GENICHIRO, et al. A61K037/20; C08B037/00.
- 
- ☐ 15. JP 06040937 A. 20 Aug 91. 15 Feb 94. ANALGESIC CONTAINING LPS AND ANALGESIC FOR ANIMAL. SOMA, GENICHIRO, et al. A61K037/20; C08B037/00.
- 
- ☐ 16. JP 05155778 A. 02 Dec 91. 22 Jun 93. GROWTH PROMOTER AND ANIMAL GROWTH PROMOTER CONTAINING LPS. SOMA, GENICHIRO, et al. A61K035/78; A61K035/66 A61K035/80 A61K037/20 A61K037/20.
- 
- ☐ 17. JP 04187640 A. 22 Nov 90. 06 Jul 92. IMMUNOLOGICAL FUNCTION PROMOTING AGENT FOR ORAL AND TRANSCUTANEOUS ADMINISTRATION AND IMMUNOLOGICAL FUNCTION PROMOTING AGENT FOR ANIMAL FOR ORAL AND TRANSCUTANEOUS ADMINISTRATION. SOMA, GENICHIRO, et al. A61K035/74; A61K035/74.
- 
- ☐ 18. JP 04099481 A. 20 Aug 90. 31 Mar 92. NOVEL BACTERIUM, NOVEL LPS, NOVEL IMMUNOFUNCTION-ACTIVATING AGENT, NEW IMMUNOFUNCTION-ACTIVATING AGENT FOR ANIMAL. SOMA, GENICHIRO, et al. 435/72 435/252.5. C12N001/20; A61K035/74 A61K039/39.
- 
- ☐ 19. JP 04049245 A. 15 Jun 90. 18 Feb 92. ANTITOXOPLASMA AGENT AND ANTITOXOPLASMA AGENT FOR ANIMAL. SOMA, GENICHIRO, et al. A61K035/78; A61K035/74 A61K035/78 A61K035/80 A61K035/84.
- 
- ☐ 20. JP 04049244 A. 15 Jun 90. 18 Feb 92. ANTIDIABETIC AGENT AND ANTIDIABETIC AGENT FOR ANIMAL. SOMA, GENICHIRO, et al. A61K035/78; A61K035/74 A61K035/78 A61K035/80 A61K035/84.
- 

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**Search Results - Record(s) 31 through 40 of 42 returned.**

- ☐ 31. JP 07126172 A TW 274517 A. Anti-MRSA drug - contains LPS and veterinary anti-MRSA drug. A61K031/715 A61K031/72 A61K035/66.
- ☐ 32. JP 07109285 A. Limulus test-positive plant lipid polysaccharide - useful as immunological activator, anti-diabetic agent, analgesic and for quasi-drug cosmetics, foods, soft drinks and feeds. A61K031/715 A61K035/78 C07G017/00 C08B037/00.
- ☐ 33. JP 06263650 A. Anticancer drug - containing LPS, used for humans and animals. A61K035/64 A61K035/72 A61K035/78 A61K035/80 A61K035/82 A61K037/20.
- ☐ 34. JP 06141849 A. LPS-producing gram-negative microbe - has high immune function activating ability for animals. A23K001/16 A61K037/20 C08B037/00 C12N001/20 C12N001/20 C12R001:01.
- ☐ 35. JP 05155778 A. Growth stimulant for animals - contains lipo:polysaccharide which activates macrophage TNF productivity. A61K035/66 A61K035/78 A61K035/80 A61K037/20.
- ☐ 36. EP 533517 A1 CA 2076505 A JP 06065092 A US 5413993 A. Lipo:polysaccharide-contg. compsns. for treating drug withdrawal symptoms - have higher therapeutic range than e.g. methadone, clonidine and dizocilpine. MIZUNO, D, et al. A61K031/715 A61K035/74 A61K037/00 A61K037/20 C08B037/00 C12N001/20 C12P019/04.
- ☐ 37. JP 04187640 A. Oral and transdermal immune function accelerating agents - contain pertussis bacteria cell lipopolysaccharide(s). A61K035/74.
- ☐ 38. EP 477050 A DE 69126183 E EP 477050 A3 EP 477050 B1. LPS-producing bacteria - has strong immunity-stimulating, analgesic and anti-withdrawal effects. MIZUNO, D, et al. A61K031/735 A61K037/20 A61K037/22 C12N001/20 C12P019/04 C12R001/01 C12N001/20 C12R001:01.
- ☐ 39. EP 472467 A CA 2049548 C CA 2049533 A CA 2049548 A JP 04099481 A EP 472467 A3 US 5281583 A JP 06040937 A JP 06078756 A JP 06090745 A US 5346891 A US 5494819 A. New analgesic compsns. for human and veterinary use - comprise lipo:polysaccharide(s) having specified macrophage activation. MIZUNO, D, et al. A61K031/70 A61K031/715 A61K035/74 A61K037/20 A61K038/16 A61K039/39 C08B037/00 C12N001/00 C12N001/20 C12P019/04 C12P021/00 C12R001/01 C12N001/20 C12R001:425 C12N001/20 C12R001:01 C12P019/04 C12R001:01 C12N001/20 C12R001:01 C12P019/04 C12R001:01.
- ☐ 40. EP 462022 A CA 2044808 A EP 462022 A3 JP 04049244 A. Human and veterinary antidiabetic agents - contain lipo:polysaccharide (LPS) from lipid A, vegetable or bacterial sources. MIZUNO, D, et al. A61K035/78 A61K037/18 A61K037/20.

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Terms	Documents
L3 and LPS	42

[Generate Collection](#)[Print](#)**Search Results - Record(s) 32 through 41 of 42 returned.**

- 
- ☐ 32. [JP 07109285 A](#). Limulus test-positive plant lipid polysaccharide - useful as immunological activator, anti-diabetic agent, analgesic and for quasi-drug cosmetics, foods, soft drinks and feeds. A61K031/715 A61K035/78 C07G017/00 C08B037/00.
- 
- ☐ 33. [JP 06263650 A](#). Anticancer drug - containing LPS, used for humans and animals. A61K035/64 A61K035/72 A61K035/78 A61K035/80 A61K035/82 A61K037/20.
- 
- ☐ 34. [JP 06141849 A](#). LPS-producing gram-negative microbe - has high immune function activating ability for animals. A23K001/16 A61K037/20 C08B037/00 C12N001/20 C12N001/20 C12R001:01.
- 
- ☐ 35. [JP 05155778 A](#). Growth stimulant for animals - contains lipo:polysaccharide which activates macrophage TNF productivity. A61K035/66 A61K035/78 A61K035/80 A61K037/20.
- 
- ☐ 36. [EP 533517 A1](#) [CA 2076505 A](#) [JP 06065092 A](#) [US 5413993 A](#). Lipo:polysaccharide-contg. compsns. for treating drug withdrawal symptoms - have higher therapeutic range than e.g. methadone, clonidine and dizocilpine. MIZUNO, D, et al. A61K031/715 A61K035/74 A61K037/00 A61K037/20 C08B037/00 C12N001/20 C12P019/04.
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- ☐ 37. [JP 04187640 A](#). Oral and transdermal immune function accelerating agents - contain pertussis bacteria cell lipopolysaccharide(s). A61K035/74.
- 
- ☐ 38. [EP 477050 A](#) [DE 69126183 E](#) [EP 477050 A3](#) [EP 477050 B1](#). LPS-producing bacteria - has strong immunity-stimulating, analgesic and anti-withdrawal effects. MIZUNO, D, et al. A61K031/735 A61K037/20 A61K037/22 C12N001/20 C12P019/04 C12R001/01 C12N001/20 C12R001:01.
- 
- ☐ 39. [EP 472467 A](#) [CA 2049548 C](#) [CA 2049533 A](#) [CA 2049548 A](#) [JP 04099481 A](#) [EP 472467 A3](#) [US 5281583 A](#) [JP 06040937 A](#) [JP 06078756 A](#) [JP 06090745 A](#) [US 5346891 A](#) [US 5494819 A](#). New analgesic compsns. for human and veterinary use - comprise lipo:polysaccharide(s) having specified macrophage activation. MIZUNO, D, et al. A61K031/70 A61K031/715 A61K035/74 A61K037/20 A61K038/16 A61K039/39 C08B037/00 C12N001/00 C12N001/20 C12P019/04 C12P021/00 C12R001/01 C12N001/20 C12R001:425 C12N001/20 C12R001:01 C12P019/04 C12R001:01 C12N001/20 C12R001:01 C12P019/04 C12R001:01.
- 
- ☐ 40. [EP 462022 A](#) [CA 2044808 A](#) [EP 462022 A3](#) [JP 04049244 A](#). Human and veterinary antidiabetic agents - contain lipo:polysaccharide (LPS) from lipid A, vegetable or bacterial sources. MIZUNO, D, et al. A61K035/78 A61K037/18 A61K037/20.
- 
- ☐ 41. [EP 462021 A](#) [CA 2044811 A](#) [EP 462021 A3](#) [JP 04049243 A](#). Human and veterinary cholesterol lowering agents - contain lipo:polysaccharide from lipid A, vegetable or bacterial sources in admixture with carrier. MIZUNO, D, et al. A61K035/78 A61K037/18 A61K037/20.
- 

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☐ 41. EP 462021 A CA 2044811 A EP 462021 A3 JP 04049243 A. Human and veterinary cholesterol lowering agents - contain lipo:polysaccharide from lipid A, vegetable or bacterial sources in admixture with carrier. MIZUNO, D, et al. A61K035/78 A61K037/18 A61K037/20.

☐ 42. EP 462020 A CA 2044802 A EP 462020 A3 JP 04049242 A. Compsn. for treating herpes - comprising at least one lipo:polysaccharide having defined macrophage activation. MIZUNO, D, et al. A61K035/78 A61K037/18 A61K037/20.

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Terms	Documents
L3 and LPS	42

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## WEST Search History

DATE: Sunday, July 20, 2003

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<i>DB=USPT,PGPB,JPAB,EPAB,DWPI; PLUR=YES; OP=OR</i>			
L6	L3 and crustacea	1	L6
L5	L3 and LPS	42	L5
L4	L3 and fish	6	L4
L3	L2 and mizuno	114	L3
L2	soma	5855	L2
L1	'MIZUNO CORPORATION!'	104	L1

END OF SEARCH HISTORY



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L5: Entry 12 of 42

File: JPAB

Apr 5, 1994

PUB-NO: JP406090745A

DOCUMENT-IDENTIFIER: JP 06090745 A

TITLE: LPS-PRODUCING MICROORGANISM, LPS AND PHARMACEUTICAL AND ANIMAL DRUG  
CONTAINING LPS

PUBN-DATE: April 5, 1994

## INVENTOR-INFORMATION:

NAME

COUNTRY

SOMA, GENICHIRO

YOSHIMURA, ATSUSHI

TSUKIOKA, DAISUKE

MIZUNO, DENICHI

OSHIMA, HARUYUKI

## ASSIGNEE-INFORMATION:

NAME

COUNTRY

CHIBA SEIFUN KK

MIZUNO DENICHISOMA GENICHIRO

APPL-NO: JP04332205

APPL-DATE: November 19, 1992

US-CL-CURRENT: 424/282.1; 435/72

INT-CL (IPC): C12N 1/20; A61K 35/74; A61K 37/20; A61K 37/20; A61K 37/20; A61K 37/20;  
C08B 37/00; C12P 19/04

## ABSTRACT:

PURPOSE: To provide a new immune function activating agent which can be administered by oral administration, transcutaneous administration and injection, an analgesic, their animal drugs, new LPS acting as the active component of the agent and new bacterial strain capable of producing the LPS.

CONSTITUTION: Three kinds of LPS compounds having the following physical properties.

(1) LPS1, principal molecular weight, 5,000±1,000 (SDS-PAGE process); number of phosphorus atoms, 2±1/5,000 molecular weight; number of hexosamine groups, 9±1/5,000 molecular weight; number of KDO groups, 2±1/5,000 molecular weight: (2) LPS2, principal molecular weight, 6,500±2,500 (SDS-PAGE process); number of phosphorus atoms, 1-2/5,000 molecular weight; number of hexosamine groups, 7±1/5,000 molecular weight; number of KDO groups, 1-2/5,000 molecular weight: (3) LPS3, principal molecular weight, 6,500±2,500 (SDS-PAGE process); number of phosphorus atoms, 2±1/5,000 molecular weight; number of hexosamine groups, 5±1/5,000 molecular weight; number of KDO groups, 2±1/5,000 molecular weight.

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L5: Entry 12 of 42

File: JPAB

Apr 5, 1994

DOCUMENT-IDENTIFIER: JP 06090745 A

TITLE: LPS-PRODUCING MICROORGANISM, LPS AND PHARMACEUTICAL AND ANIMAL DRUG  
CONTAINING LPSAbstract Text (1):

PURPOSE: To provide a new immune function activating agent which can be administered by oral administration, transcutaneous administration and injection, an analgesic, their animal drugs, new LPS acting as the active component of the agent and new bacterial strain capable of producing the LPS.

Abstract Text (2):

CONSTITUTION: Three kinds of LPS compounds having the following physical properties.

(1) LPS1, principal molecular weight,  $5,000 \pm 1,000$  (SDS-PAGE process); number of phosphorus atoms,  $2 \pm 1/5,000$  molecular weight; number of hexosamine groups,  $9 \pm 1/5,000$  molecular weight; number of KDO groups,  $2 \pm 1/5,000$  molecular weight: (2) LPS2, principal molecular weight,  $6,500 \pm 2,500$  (SDS-PAGE process); number of phosphorus atoms,  $1-2/5,000$  molecular weight; number of hexosamine groups,  $7 \pm 1/5,000$  molecular weight; number of KDO groups,  $1-2/5,000$  molecular weight: (3) LPS3, principal molecular weight,  $6,500 \pm 2,500$  (SDS-PAGE process); number of phosphorus atoms,  $2 \pm 1/5,000$  molecular weight; number of hexosamine groups,  $5 \pm 1/5,000$  molecular weight; number of KDO groups,  $2 \pm 1/5,000$  molecular weight.

Applicant Name (2):MIZUNO DENICHIApplicant Name (3):SOMA GENICHIROApplicant Name (Derived) (2):MIZUNO DENICHIApplicant Name (Derived) (3):SOMA GENICHIROInventor Name (1):SOMA, GENICHIROInventor Name (4):MIZUNO, DENICHIInventor Name ( Derived ) (1):SOMA, GENICHIROInventor Name ( Derived ) (4):MIZUNO, DENICHI

☐ Generate Collection

L5: Entry 16 of 42

File: JPAB

Jun 22, 1993

PUB-NO: JP405155778A

DOCUMENT-IDENTIFIER: JP 05155778 A

TITLE: GROWTH PROMOTER AND ANIMAL GROWTH PROMOTER CONTAINING LPS

PUBN-DATE: June 22, 1993

## INVENTOR-INFORMATION:

NAME

COUNTRY

SOMA, GENICHIRO

YOSHIMURA, ATSUSHI

TSUKIOKA, DAISUKE

MIZUNO, DENICHI

OSHIMA, HARUYUKI

INT-CL (IPC): A61K 35/78; A61K 35/66; A61K 35/80; A61K 37/20; A61K 37/20

## ABSTRACT:

PURPOSE: To obtain a growth promoter and an animal growth promoter, having high growth promoting effects, usable for a long period and administrable by any of oral or percutaneous route, transcutan bath or injection.

CONSTITUTION: A growth promoter is characterized by including at least one of the following lipopolysaccharide (LPS) therein: An LPS capable of providing a content of the LPS, positive to the Limulus test and affording an ED50 of the macrophage activating ability of 0.4-100ng/ml culture solution when sigmoid curves are drawn by using the macrophage activating ability of the LPS activating the TNF productivity of an in vitro cultured macrophage as an index and plotting the macrophage activating ability (%) in the macrophage activating ability providing the TNF production of the macrophage without adding the LPS as 0%, and the macrophage activating ability when the TNF production of the macrophage attains the maximum constant amount as 100% on the coordinate axis versus the content of the LPS positive to the Limulus test at that time expressed on the logarithmic scale on the abscissa axis.

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L5: Entry 17 of 42

File: JPAB

Jul 6, 1992

PUB-NO: JP404187640A

DOCUMENT-IDENTIFIER: JP 04187640 A

TITLE: IMMUNOLOGICAL FUNCTION PROMOTING AGENT FOR ORAL AND TRANSCUTANEOUS  
ADMINISTRATION AND IMMUNOLOGICAL FUNCTION PROMOTING AGENT FOR ANIMAL FOR ORAL AND  
TRANSCUTANEOUS ADMINISTRATION

PUBN-DATE: July 6, 1992

## INVENTOR-INFORMATION:

NAME

COUNTRY

SOMA, GENICHIRO

YOSHIMURA, ATSUSHI

TSUKIOKA, DAISUKE

MIZUNO, DENICHI

OSHIMA, HARUYUKI

## ASSIGNEE-INFORMATION:

NAME

COUNTRY

CHIBA SEIFUN KK

MIZUNO DENICHISOMA GENICHIRO

APPL-NO: JP02315919

APPL-DATE: November 22, 1990

INT-CL (IPC): A61K 35/74; A61K 35/74

## ABSTRACT:

PURPOSE: To provide an immunological function promoting agent for oral and transcutaneous administration, containing Bordetella pertussis LPS, having high immunological function activation capability and high chemotherapeutic index and producible at a low cost.

CONSTITUTION: The objective agent contains Bordetella pertussis LPS preferably having a molecular weight of 6000±1000 and 9000±1000 (determined by SDS electrophoresis) and containing 5 P atoms, 16±2 hexosamine groups, 5 fatty acid groups and 2±1 KDO based on 8,000 molecular weight. The immunological function is e.g. macrophage-activation property, especially endogenous TNF- production promoting property. The immunological function promoting activity can be further improved by the combined use of an agent for suppressing the production of prostaglandin (e.g. indomethacin).

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L5: Entry 18 of 42

File: JPAB

Mar 31, 1992

PUB-NO: JP404099481A

DOCUMENT-IDENTIFIER: JP 04099481 A

TITLE: NOVEL BACTERIUM, NOVEL LPS, NOVEL IMMUNOFUNCTION-ACTIVATING AGENT, NEW  
IMMUNOFUNCTION-ACTIVATING AGENT FOR ANIMAL

PUBN-DATE: March 31, 1992

## INVENTOR-INFORMATION:

NAME

COUNTRY

SOMA, GENICHIRO

YOSHIMURA, ATSUSHI

TSUKIOKA, DAISUKE

MIZUNO, DENICHI

OSHIMA, HARUYUKI

US-CL-CURRENT: 435/72; 435/252.5

INT-CL (IPC): C12N 1/20; A61K 35/74; A61K 39/39

## ABSTRACT:

NEW MATERIAL: A LPS-producing gram negative short Bacillus bacterium (FERN P-3509).

Shape: a short culmed shape, not moving and negative to Gram stain; growth state:

forms a yellow-creamy round and opaque colony in the standard agar medium;

physiological properties: positive to Voges-Proskauer reaction, O-F test, etc., and

negative to indole-producing reaction. etc.; the utilization of carbon sources:

utilizes lactose, rhamnose, etc., not utilize adonite, inositol, etc.

USE: A bacterium for producing a novel LPS which is an active ingredient for  
immunofunction-activating agents for animals.PREPARATION: Wheat flour is mixed with distilled water, cultured with shaking at  
37°C, diluted, sowed on a standard agar medium and subsequently cultured. A colony  
producing the LPS is screened from the colonies thus produced by a test.

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L5: Entry 30 of 42

File: DWPI

Aug 1, 1996

DERWENT-ACC-NO: 1996-362635

DERWENT-WEEK: 199636

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TITLE: Low mol. wt. lipo-polysaccharide from Pantoea or Salmonella - has high safety and high biological activity for pharmaceutical use, e.g. against toxoplasmosis and hypocholesterolaemia, anticancer and antidiabetic agent

INVENTOR: MIZUNO, D; NISHIZAWA, T ; SOMA, G

PATENT-ASSIGNEE:

ASSIGNEE

MIZUNO DSOMA G

TAIHO PHARM CO LTD

CODE

MIZUI

SOMAI

TAIH

PRIORITY-DATA: 1995JP-0012126 (January 27, 1995)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
WO 9623002 A1	August 1, 1996	J	032	C08B037/00
JP 08198902 A	August 6, 1996		011	C08B037/00

DESIGNATED-STATES: US AT BE CH DE DK ES FR GB GR IE IT LU MC NL PT SE

CITED-DOCUMENTS: JP 40187640; JP 5155778 ; JP 60141849

APPLICATION-DATA:

PUB-NO	APPL-DATE	APPL-NO	DESCRIPTOR
WO 9623002A1	January 25, 1996	1996WO-JP00135	
JP 08198902A	January 27, 1995	1995JP-0012126	

INT-CL (IPC): A61 K 31/715; C08 B 37/00; C12 P 19/04; C12 P 19/04; C12 R 1:42; C12 P 19/04; C12 R 1:01; C12 P 19/04; C12 R 1:42

ABSTRACTED-PUB-NO: WO 9623002A

BASIC-ABSTRACT:

Low mol. wt. lipopolysaccharide of microbial origin (LP) has the following properties: (a) mol.wt. 5000 (plus or minus 2000) by SDS-PAGE using protein markers, the material being substantially free from fractions of other mol.wt.; (b) 1-3 hexosamine units/mol. (5000 mol.wt. by Elson-Morgan method); (c) 1-3 2keto-3-deoxy-octonate units/mol. (5000 mol.wt. by the diphenylamine method); (d) limulus activity of 10 EU/ng, and (e) protein content and nucleic acid content each <1% (wt.).

USE - (LP) can be used as a pharmaceutical, e.g. against toxoplasmosis, as hypocholesterolaemia, anti-herpes, anti-rheumatic, anti-diabetic, immuno-modulator, anti-cancer, anti-gastric ulcer or anti-haemorrhoid agent. (LP) should be administered at 1 mug-100 mg orally or 10 ng-10 mg parenterally/day.

ADVANTAGE - (LP) has a high safety factor (low toxicity) and high biological activity.

CHOSEN-DRAWING: Dwg.0/2

TITLE-TERMS: LOW MOLECULAR WEIGHT LIPO POLYSACCHARIDE SALMONELLA HIGH SAFETY HIGH  
BIOLOGICAL ACTIVE PHARMACEUTICAL TOXOPLASMA HYPOCHOLESTEROLAEMIC ANTICANCER  
ANTIDIABETIC AGENT

DERWENT-CLASS: B04 D16

CPI-CODES: B04-C02V; B14-A02A3; B14-C06; B14-D02A2; B14-E04; B14-E08; B14-G03;  
B14-S04; D05-C; D05-H13;

CHEMICAL-CODES:

Chemical Indexing M1 \*01\*

Fragmentation Code

M423 M710 M903 P210 P220 P423 P433 P434 P529 P633

P738 P816 Q233 V735

SECONDARY-ACC-NO:

CPI Secondary Accession Numbers: C1996-114244





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L5: Entry 34 of 42

File: DWPI

May 24, 1994

DERWENT-ACC-NO: 1994-205014

DERWENT-WEEK: 199425

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TITLE: LPS-producing gram-negative microbe - has high immune function activating ability for animals

Basic Abstract Text (1):

An LPS-producing Gram-negative microbe has the following properties: (a) Form (1) Motile at 30 deg.C, (2) Gram-staining:- (b) Growing condition (1) Trypticase soya agar medium: Forms white to pale yellow colonies. (c) Physiological properties (1) Indol formation:+; (2) Voges-Proskauer reaction:+; (3) Nitrate reduction:+; (4) hydrogen sulphide formation:+/-; (5) Esculin hydrolysis:+; (6) Gas formation from D-glucose:+; (7) Lysine decarboxylase: +/-; (8) Arginine dihydrolyase:+; (9) Ornithine decarboxylase:-; (10) Phenylalanine deaminase:-; (11) Deoxyribonuclease:+; (12) Kovac oxidase:+; (13) Beta-galactosidase:+; (14) Urease:-; (d) Utility (1) Simmons citrate: +/-; (2) Malonic acid:-; (3) Inositol:-; (4) Sucrose: +/-; (5) Adonitol:-; (6) Raffinose:-; (7) L-arabinose:+; (8) D-mannitol:+/-; (9) L-rhamnose:-; (10) D-sorbitol:-.

Basic Abstract Text (2):

LPS originated from the microbe has the following properties: M.W.: 5000+/-1500 (by tricine SDS-PAGE); Phosphor molar number: 2.0+/-1/M.W.5000; Hexosamine molar number: 9.1+/-M.W.5000; KDO molar number: 0.8+/-0.5/M.W.5000.

Basic Abstract Text (3):

An immune function activator contg. the LPS and an immune function activator for animals contg. above LPS, are also claimed.

Basic Abstract Text (4):

USE/ADVANTAGE - The LPS is new and low in toxicity, has a high immune function activating ability, can be dosed through various paths and can be produced at low cost.

Basic Abstract Text (5):

In an example, E coli LPS0127:BS was cultured in a trypticase soya agar medium at 30 deg.C overnight. Among 7 colonies, a white to pale yellow Gram-negative colony was collected and cultured in an LB medium at 30 deg.C for 18 hrs. The culture was centrifuged and the ppt. was suspended in distilled water, treated with phenol, centrifuged and extracted with distilled water. The extract was dialysed against distilled water, purified by a Q-Sepharose column and eluted by 400 mM NaCl, ultrafiltered and freeze-dried to give 13.0 mg LPS. It showed a TNF productivity 2.5 times higher than that of E coli LPS.

Patent Assignee Terms (1):MIZUNO DPatent Assignee Terms (2):SOMA GPatent Assignee Terms (1):MIZUNO DPatent Assignee Terms (2):SOMA G

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L5: Entry 35 of 42

File: DWPI

Jun 22, 1993

DERWENT-ACC-NO: 1993-232272

DERWENT-WEEK: 199329

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TITLE: Growth stimulant for animals - contains lipo:polysaccharide which activates macrophage TNF productivity

PATENT-ASSIGNEE: CHIBA SEIFUN KK (CHIBN), MIZUNO D (MIZUI), SOMA G (SOMAI)

PRIORITY-DATA: 1991JP-0357351 (December 2, 1991)

## PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
JP 05155778 A	June 22, 1993		031	A61K035/78

## APPLICATION-DATA:

PUB-NO	APPL-DATE	APPL-NO	DESCRIPTOR
JP 05155778A	December 2, 1991	1991JP-0357351	

INT-CL (IPC): A61K 35/66; A61K 35/78; A61K 35/80; A61K 37/20

ABSTRACTED-PUB-NO: JP 05155778A

## BASIC-ABSTRACT:

Growth stimulants contain at least one lipopolysaccharide (LPS) which activates the macrophage TNF productivity obtd. by in vitro culture as an index, and contains Limulus test positive LPS exhibiting ED50 for macrophage activity or 0.4-100 ng/ml of LPS in culture soln. expressed on a graph having ordinate axis of TNF productivity of macrophage placing macrophage activation in the absence of LPS at 0% and macrophage activation to make max. macrophage TNF prodn. at 100%, and a transverse axis expressing Limulus test positive LPS content in logarithmic scale.

Also claimed are (1) growth stimulant LPS obtd. from plants, bacteria, lipid A, and their mixts.; (2) a growth stimulant LPS obtd. from wheat and having following physicochemical properties: (a) main molecular wt. of 8,000 +/- 1,000 (SDS-1 method), 8,000 +/- 1,000 (SDS-2 method), and 5,000 +/- 2,000 (SDS-2 method), amt. of P is 1-4/molecular wt. 8,000 amt. of hexosamine: 6 +/- 2/molecular wt. 8,000, amt. of fatty acid: 6 +/- 2/molecular wt. 8,000, and KDO (2-keto-3-deoxyoctonate) number is 5 +/- 1/molecular wt. 8,000; (3) growth stimulant LPS obtd. from chlorella, E. coli, Bordetella pertussis, Agrobacterium radiobacter and three bacteria having similar physicochemical properties are also claimed.

USE/ADVANTAGE - Growth stimulant for animals including humans for the prevention of premature infants without adverse reactions for in vivo residual activity which can be administered for a long period by oral, percutaneous, medicated bath or injection and can be massively produced at low cost

ABSTRACTED-PUB-NO: JP 05155778A

## EQUIVALENT-ABSTRACTS:

CHOSEN-DRAWING: Dwg.0/0

DERWENT-CLASS: B04 C03

CPI-CODES: B04-B01B; C04-B01B; B04-B04J; C04-B04J; B04-C02; C04-C02;



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L5: Entry 35 of 42

File: DWPI

Jun 22, 1993

DERWENT-ACC-NO: 1993-232272

DERWENT-WEEK: 199329

COPYRIGHT 2003 DERWENT INFORMATION LTD

TITLE: Growth stimulant for animals - contains lipo:polysaccharide which activates macrophage TNF productivity

Basic Abstract Text (1):

Growth stimulants contain at least one lipopolysaccharide (LPS) which activates the macrophage TNF productivity obtd. by in vitro culture as an index, and contains Limulus test positive LPS exhibiting ED50 for macrophage activity or 0.4-100 ng/ml of LPS in culture soln. expressed on a graph having ordinate axis of TNF productivity of macrophage placing macrophage activation in the absence of LPS at 0% and macrophage activation to make max. macrophage TNF prodn. at 100%, and a transverse axis expressing Limulus test positive LPS content in logarithmic scale.

Basic Abstract Text (2):

Also claimed are (1) growth stimulant LPS obtd. from plants, bacteria, lipid A, and their mixts.; (2) a growth stimulant LPS obtd. from wheat and having following physicochemical properties: (a) main molecular wt. of 8,000 +/- 1,000 (SDS-1 method), 8,000 +/- 1,000 (SDS-2 method), and 5,000 +/- 2,000 (SDS-2 method), amt. of P is 1-4/molecular wt. 8,000 amt. of hexosamine: 6 +/- 2/molecular wt. 8,000, amt. of fatty acid: 6 +/- 2/molecular wt. 8,000, and KDO (2-keto-3-deoxyoctonate) number is 5 +/- 1/molecular wt. 8,000; (3) growth stimulant LPS obtd. from chlorella, E. coli, Bordetella pertussis, Agrobacterium radiobacter and three bacteria having similar physicochemical properties are also claimed.

Patent Assignee Terms (2):MIZUNO DPatent Assignee Terms (3):SOMA GPatent Assignee Terms (2):MIZUNO DPatent Assignee Terms (3):SOMA G

☐ Generate Collection

L5: Entry 38 of 42

File: DWPI

Mar 25, 1992

DERWENT-ACC-NO: 1992-098698

DERWENT-WEEK: 199731

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TITLE: LPS-producing bacteria - has strong immunity-stimulating, analgesic and anti-withdrawal effects

Basic Abstract Text (1):

Three new strains of LPS-producing gram-negative small bacilli have morphological, growth, physiological and C utilisation characteristics defined in the specification.

Basic Abstract Text (2):

Also claimed are: (1) LPS provided by the 1st strain of bacilli (LPS1) having a dominant mol.wt. of 5,000 +/- 1,000 (SDS-PAGE), having 2+/-1 phosphorus, 9+/-1 hexosamines and 2+/-1 KDO per mol.wt. of 5,000; (2) LPS provided by the 2nd strain of bacilli (LPS2) having a dominant mol.wt. of 6,500+/-2,500 (SDS-PAGE), having 1-2 phosphorus, 7+/-1 hexosamines and 1-2 KDO per mol.wt. of 5,000; (3) LPS provided by the 3rd strain of bacilli (LPS3) having a dominant mol.wt. of 6,500 +/- 2,500 (SDS-PAGE), having 2+/-1 phosphorus, 5+/-1 hexosamines and 2+/-1 KDO per mol.wt. of 5,000; and (4) immunity stimulators, analgesics and antiwithdrawal agents contg. LPS as above.

Patent Assignee Terms (1):MIZUNO DPatent Assignee Terms (2):SOMA GPatent Assignee Terms (1):MIZUNO DPatent Assignee Terms (2):SOMA GEquivalent Abstract Text (1):

LPS (lipopolysaccharide)-producing gram-negative small bacilli which have the intrinsic characteristics of the microorganism FERM BP-3509 belonging to the genus Serratia, the characteristics being as follows: (a) Morphological characteristics (1) Small rod, (2) No motility; (3) Gram stain; (b) Growth (1) Standard agar medium: a yellow to creamy round opaque colony is formed, (2) SS (Salmonella-Shigella) agar medium: A white translucent colony is formed, (3) TSI (Triple Sugar iron) medium: No change is found on the slant, but the higher layer changes to yellow and Gas is produced; (c) Physiological characteristics (1) Voges-Proskauer reaction + (2) Indole production - (3) Hydrogen sulphide production - (4) utilisation of citrate + (5) Urease - (6) Oxidase - (7) O-F test (Oxidation-Fermentation test) + (d) Utilisation of carbon sources (1) Lactose + (2) Adonitol - (3) Rhamnose + (4) Mannitol + (5) Esculin + (6) Inositol - (7) Sorbitol + (8) Arabinose + (9) Raffinose + (10) Sucrose + (e) Others (1) Lysin decarboxylase - (2) Utilisation of malonate - (3) Arginine dihydroxylase - (4) Phenylalanine deaminase - (5) Omithine decarboxylase.

Inventor Name (1):MIZUNO, DInventor Name (3):SOMA, G

•  
•  
Inventor Name (6) :  
SOMA, G I



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L5: Entry 38 of 42

File: DWPI

Mar 25, 1992

DERWENT-ACC-NO: 1992-098698

DERWENT-WEEK: 199731

COPYRIGHT 2003 DERWENT INFORMATION LTD

TITLE: LPS-producing bacteria - has strong immunity-stimulating, analgesic and anti-withdrawal effects

INVENTOR: MIZUNO, D; OSHIMA, H ; SOMA, G ; TSUKIOKA, D ; YOSHIMURA, K ; SOMA, G IPATENT-ASSIGNEE: MIZUNO D (MIZUI), SOMA G (SOMAI), CHIBA FLOUR MILLING CO LTD (CHIBN)

PRIORITY-DATA: 1990JP-0312932 (November 20, 1990), 1990JP-0218599 (August 20, 1990)

## PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
EP 477050 A	March 25, 1992		027	
DE 69126183 E	June 26, 1997		000	C12N001/20
EP 477050 A3	March 17, 1993		000	
EP 477050 B1	May 21, 1997	E	031	C12N001/20

DESIGNATED-STATES: AT BE CH DE DK ES FR GB GR IT LI LU NL SE AT BE CH DE DK ES FR GB GR IT LI LU NL SE

CITED-DOCUMENTS: No-SR.Pub; 9.Jnl.Ref ; 5.Jnl.Ref ; EP 472467 ; EP 533517

## APPLICATION-DATA:

PUB-NO	APPL-DATE	APPL-NO	DESCRIPTOR
EP 477050A	August 20, 1991	1991EP-0402275	
DE 69126183E	August 20, 1991	1991DE-0626183	
DE 69126183E	August 20, 1991	1991EP-0402275	
DE 69126183E		EP 477050	Based on
EP 477050A3	August 20, 1991	1991EP-0402275	
EP 477050B1	August 20, 1991	1991EP-0402275	

INT-CL (IPC): A61K 31/735; A61K 37/20; A61K 37/22; C12N 1/20; C12P 19/04; C12R 1/01; C12N 1/20; C12R 1/01

RELATED-ACC-NO: 1992-066655; 1993-095774

ABSTRACTED-PUB-NO: EP 477050A

## BASIC-ABSTRACT:

Three new strains of LPS-producing gram-negative small bacilli have morphological, growth, physiological and C utilisation characteristics defined in the specification.

Also claimed are: (1) LPS provided by the 1st strain of bacilli (LPS1) having a dominant mol.wt. of 5,000 +/- 1,000 (SDS-PAGE), having 2+/-1 phosphorus, 9+/-1 hexosamines and 2+/-1 KDO per mol.wt. of 5,000; (2) LPS provided by the 2nd strain of bacillis (LPS2) having a dominant mol.wt. of 6,500+/-2,500 (SDS-PAGE), having 1-2 phosphorus, 7+/-1 hexosamines and 1-2 KDO per mol.wt. of 5,000; (3) LPS provided by the 3rd strain of bacilli (LPS3) having a dominant mol.wt. of 6,500 +/- 2,500 (SDS-PAGE), having 2+/-1 phosphorus, 5+/-1 hexosamines and 2+/-1 KDO per mol.wt. of

5,000; and (4) immunity stimulators, analgesics and antiwithdrawal agents contg. LPS as above.

USE/ADVANTAGE - The LPSs show excellent immunity-stimulating, analgesic and antiwithdrawal effects and a high therapeutic range. They exhibit none of the side effects and drawbacks of prior art agents (e.g. dependency) and can be produced in large amts. at low cost. They can be used to treat both human and animal patients. For human administration, dose is pref. 1ng- 100mg (oral), 10ng-10mg (i.v.) and 100ng-1mg (percutaneous) per day (60kg adult). For veterinary use, corresponding values are given depending on body wt.

ABSTRACTED-PUB-NO: EP 477050B

EQUIVALENT-ABSTRACTS:

LPS (lipopolysaccharide)-producing gram-negative small bacilli which have the intrinsic characteristics of the microorganism FERM BP-3509 belonging to the genus Serratia, the characteristics being as follows: (a) Morphological characteristics (1) Small rod, (2) No motility, (3) Gram stain; (b) Growth (1) Standard agar medium: a yellow to creamy round opaque colony is formed, (2) SS (Salmonella-Shigella) agar medium: A white translucent colony is formed, (3) TSI (Triple Sugar iron) medium: No change is found on the slant, but the higher layer changes to yellow and Gas is produced; (c) Physiological characteristics (1) Voges-Proskauer reaction + (2) Indole production - (3) Hydrogen sulphide production - (4) utilisation of citrate + (5) Urease - (6) Oxidase - (7) O-F test (Oxidation-Fermentation test) + (d) Utilisation of carbon sources (1) Lactose + (2) Adonitol - (3) Rhamnose + (4) Mannitol + (5) Esculin + (6) Inositol - (7) Sorbitol + (8) Arabinose + (9) Raffinose + (10) Sucrose + (e) Others (1) Lysin decarboxylase - (2) Utilisation of malonate - (3) Arginine dihydroxylase - (4) Phenylalanine deaminase - (5) Omithine decarboxylase.

CHOSEN-DRAWING: Dwg.0/5 Dwg.0/5

DERWENT-CLASS: B04 C06 D16

CPI-CODES: B04-B01B; C04-B01B; B04-B02B1; C04-B02B1; B04-C02F; C04-C02F; B12-A01; C12-A01; B12-A06; C12-A06; B12-D01; C12-D01; B12-J05; C12-J05; D05-C08; D05-H04;



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L5: Entry 39 of 42

File: DWPI

Feb 26, 1992

DERWENT-ACC-NO: 1992-066655

DERWENT-WEEK: 200253

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TITLE: New analgesic compsns. for human and veterinary use - comprise  
lipo:polysaccharide(s) having specified macrophage activation

INVENTOR: MIZUNO, D; OSHIMA, H ; SOMA, G ; TSUKIOKA, D ; YOSHIMURA, K ; SOMA, G IPATENT-ASSIGNEE: SOMA G I (SOMAI), CHIBA FLOUR MILLING CO LTD (CHIBN), MIZUNO D (MIZUI), SOMA G (SOMAI), CHIBA SEIFUN KK (CHIBN)

PRIORITY-DATA: 1990JP-0312932 (November 20, 1990), 1990JP-0218599 (August 20, 1990)

## PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
EP 472467 A	February 26, 1992		000	
CA 2049548 C	July 2, 2002	E	000	C12N001/20
CA 2049533 A	February 21, 1992		000	
CA 2049548 A	February 21, 1992		000	
JP 04099481 A	March 31, 1992		014	
EP 472467 A3	March 17, 1993		000	
US 5281583 A	January 25, 1994		025	A61K031/70
JP 06040937 A	February 15, 1994		031	A61K037/20
JP 06078756 A	March 22, 1994		016	C12N001/20
JP 06090745 A	April 5, 1994		013	C12N001/20
US 5346891 A	September 13, 1994		014	A61K031/715
US 5494819 A	February 27, 1996		013	C12N001/00

DESIGNATED-STATES: AT BE CH DE ES FR GB GR IT LI LU NL SE

CITED-DOCUMENTS: NoSR.Pub; 6.Jnl.Ref ; EP 462020 ; EP 462021 ; EP 462022

## APPLICATION-DATA:

PUB-NO	APPL-DATE	APPL-NO	DESCRIPTOR
EP 472467A	August 20, 1991	1991EP-0402276	
CA 2049548C	August 20, 1991	1991CA-2049548	
JP 04099481A	August 20, 1990	1990JP-0218599	
EP 472467A3	August 20, 1991	1991EP-0402276	
US 5281583A	August 20, 1991	1991US-0747728	CIP of
US 5281583A	August 22, 1991	1991US-0748808	
JP 06040937A	August 20, 1991	1991JP-0291844	
JP 06078756A	August 20, 1991	1991JP-0291845	
JP 06090745A	August 20, 1991	1991JP-0291845	Div ex
JP 06090745A	August 20, 1991	1992JP-0332205	
US 5346891A	August 20, 1991	1991US-0747633	
US 5494819A	August 20, 1991	1991US-0747633	Div ex
US 5494819A	April 12, 1994	1994US-0226636	
US 5494819A		US 5346891	Div ex



INT-CL (IPC): A61K 31/70; A61K 31/715; A61K 35/74; A61K 37/20; A61K 38/16; A61K 39/39; C08B 37/00; C12N 1/00; C12N 1/20 ; C12P 19/04; C12P 21/00; C12R 1/01; C12N 1/20; C12R 1/425; C12N 1/20; C12R 1/01; C12P 19/04; C12R 1/01; C12N 1/20; C12R 1/01; C12P 19/04; C12R 1/01

RELATED-ACC-NO: 1992-098698;1993-095774

ABSTRACTED-PUB-NO: EP 472467A  
BASIC-ABSTRACT:

Analgesic compsn. contains at least one lipopolysaccharide (LPS) having ED50 for macrophage activation 0.4-100 ng/ml culture, expressed in terms of limulus-test positive LPS content.

The ED50 is determined from a sigmoid plot of the macrophage activating ability (MAA; ordinate; i.e. the ability of LPS to activate prodn. of tumour necrosis factor (TNF) by macrophages cultured in vitro) vs. LPS content (abscissa). MAA is defined as 0% where no LPS is added and 100% is the max. (plateau) value. LPS is formulated with a pharmaceutically or veterinarily acceptable carrier. Specifically LPS is of plant or bacterial origin, or is synthetic lipid A.

USE/ADVANTAGE - The compsns. are useful in human or veterinary medicine; have a high therapeutic range; are inexpensive; can be prepd. on a large scale, and can be admin. orally, intradermally or by injection. The compsns. are also said to have cholesterol-lowering activity (not claimed).

ABSTRACTED-PUB-NO: US 5281583A  
EQUIVALENT-ABSTRACTS:

Analgesic compsn. comprises a macrophage-activating lipo- polysaccharide (e.g. vegetable or bacterial lipopolysaccharide or lipid A) having ED50 0.4-100 ng/cm<sup>3</sup> of culture medium and exhibiting a positive limulus test.

USE/ADVANTAGE - The prods. are administered orally, intradermally or by injection for the prophylaxis or alleviation of pain in humans or animals. The prods. are inexpensive, non-addictive and without side effects.

US 5346891A

Lipopolysaccharide produced by *Serratia ficaria* FERM BP-3509 has a mol.wt. of 5,000 +/- 1,000 by SDS-PAGE, has 1-3 phosphorous, 8-10 hexoamines and 1-3 KDO per mol. wt. of 5,000. Further characteristics are also claimed.

USE - As an immunostimulant, analgesic and antiwithdrawal agent.

US 5494819A

A biologically pure culture of a strain of the species *Pantoea agglomerans* having all the identifying characteristics of FERM BP-3511 wherein the identifying characteristics are as follows:

a) Morphological characteristics: 1) Small rod, 2) No Motility,

3) Gram stain; b) Growth: 1) Standard agar medium: A yellow round translucent colony is formed, 2) SS agar medium: No colony is formed, 3) TSI agar medium: No change is found on the slant, but the higher layer changes to yellow; Gas is not produced; c) Physiological characteristics: 1) Voges-Proskauer reaction (+); 2) Indole production (-), 3) Hydrogen sulfide production (-), 4) Utilization of citrate (+), 5) Urease (-), 6) Oxidase (-), 7) O-F test (+); d) Utilization of carbon sources: 1) Lactose (+), 2) Adonitol (-), 3) Rhamnose (+), 4) Mannitol (+), 5) Esculin (+), 6) Inositol (-), 7) Sorbitol (+), 8) Arabinose (+), 9) Raffinose (-), 10) Sucrose (+); e) Others: 1) Lysin decarboxylase (-), 2) Utilization of malonate (+), 3) Arginine dihydroxylase (-), 4) Phenylalanine deaminase (-), 5) Ornithine decarboxylase (-), f) and which produces a lipopolysaccharide having a dominant molecular weight of 6,500-/+2,500 as determined by SDS-PAGE method, 2-/+1 phosphorus, 5-/+1 hexosamines

and 2-/ +1 2-keto-3-deoxyoctonate per molecular weight of 5,000.

CHOSEN-DRAWING: Dwg.0/1 Dwg.0/0 Dwg.0/3 Dwg.0/5

DERWENT-CLASS: B04 C03

CPI-CODES: B04-B01B; B04-C02D; B04-C02F; B12-D01; B12-H03; C04-B01B; C04-C02D;  
C04-C02F; C12-D01; C12-H03;

**WEST**[Generate Collection](#)**Search Results - Record(s) 1 through 10 of 19 returned.**☐ 1. Document ID: US 6015910 A

L13: Entry 1 of 19 File: USPT Jan 18, 2000

US-PAT-NO: 6015910

DOCUMENT-IDENTIFIER: US 6015910 A

TITLE: Intermediates to pesticidal  
5-amino-4-ethylsulfinyl-1-arylpyrazoles

DATE-ISSUED: January 18, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Wu; Tai-Teh	Chapel Hill	NC	N/A	N/A

US-CL-CURRENT: 548/367.7; 548/367.4; 548/371.7

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 2. Document ID: US 5972642 A

L13: Entry 2 of 19 File: USPT Oct 26, 1999

US-PAT-NO: 5972642

DOCUMENT-IDENTIFIER: US 5972642 A

TITLE: Astaxanthin-producing yeast cells, methods for their  
preparation and their use

DATE-ISSUED: October 26, 1999

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP	CODE	COUNTRY
Flen.o slashed.; Bent	Stenl.o slashed.se	N/A	N/A		DKX
Christensen; Ib	Aller.o slashed.d	N/A	N/A		DKX
Larsen; Robert	Virum	N/A	N/A		DKX
Johansen; Steffen	Ega	N/A	N/A		DKX
Radich					
Johnson; Eric A.	Madison	WI	N/A		N/A

US-CL-CURRENT: 435/67; 435/171; 435/255.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 3. Document ID: US 5883112 A

L13: Entry 3 of 19 File: USPT Mar 16, 1999

US-PAT-NO: 5883112

DOCUMENT-IDENTIFIER: US 5883112 A

TITLE: Synergistic compositions comprising pesticidal  
5-amino-4-ethylsulfinyl-1-arylpyrazoles and piperonyl butoxide

DATE-ISSUED: March 16, 1999

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Pilato; Michael Thomas	Cary	NC	N/A	N/A
Haas; Charles Lee	Garner	NC	N/A	N/A

US-CL-CURRENT: 514/404; 514/407, 548/367.4, 548/367.7, 548/371.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 4. Document ID: US 5814652 A

L13: Entry 4 of 19 File: USPT Sep 29, 1998

US-PAT-NO: 5814652

DOCUMENT-IDENTIFIER: US 5814652 A

TITLE: Pesticidal 5-amino-4-ethylsulfinyl-1-arylpyrazoles

DATE-ISSUED: September 29, 1998

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Wu; Tai-Teh	Chapel Hill	NC	N/A	N/A

US-CL-CURRENT: 514/404; 548/367.4

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 5. Document ID: US 5712110 A

L13: Entry 5 of 19 File: USPT Jan 27, 1998

US-PAT-NO: 5712110  
DOCUMENT-IDENTIFIER: US 5712110 A

TITLE: Astaxanthin-producing yeast cells methods for their  
preparation and their use

DATE-ISSUED: January 27, 1998

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP	CODE	COUNTRY
Flen.o slashed. ; Bent	Stenl.o slashed.se	N/A	N/A		DKX
Christensen; Ib	Aller.o slashed.d	N/A	N/A		DKX
Larsen; Robert	Virum	N/A	N/A		DKX
Johansen; Steffen	Ega	N/A	N/A		DKX
Radich					
Johnson; Eric A.	Madison	WI	N/A		N/A

US-CL-CURRENT: 435/67; 435/171, 435/255.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 6. Document ID: US 5709856 A

L13: Entry 6 of 19 File: USPT Jan 20, 1998

US-PAT-NO: 5709856  
DOCUMENT-IDENTIFIER: US 5709856 A

TITLE: Astaxanthin-producing yeast cells, methods for their  
preparation and their use

DATE-ISSUED: January 20, 1998

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP	CODE	COUNTRY
Flen.o slashed.; Bent	Stenl.o slashed.se	N/A	N/A		DKX
Christensen; Ib	Aller.o slashed.d	N/A	N/A		DKX
Larsen; Robert	Virum	N/A	N/A		DKX
Johansen; Steffen	Ega	N/A	N/A		DKX
Radich					
Johnson; Eric A.	Madison	WI	N/A		N/A

US-CL-CURRENT: 424/93.51; 426/540, 426/62, 435/255.1, 435/67

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 7. Document ID: US 5698246 A

L13: Entry 7 of 19

File: USPT

Dec 16, 1997

US-PAT-NO: 5698246

DOCUMENT-IDENTIFIER: US 5698246 A

TITLE: Foodstuff for and method of feeding crustaceans and fish

DATE-ISSUED: December 16, 1997

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Villamar; Daniel F.	Maple Grove	MN	N/A	N/A

US-CL-CURRENT: 426/54; 426/2, 426/53, 426/623, 426/635

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KVMC	Draw. Desc	Image
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☐ 8. Document ID: US 5679567 A

L13: Entry 8 of 19

File: USPT

Oct 21, 1997

US-PAT-NO: 5679567

DOCUMENT-IDENTIFIER: US 5679567 A

TITLE: Astaxanthin-producing yeast cells, methods for their preparation and their use

DATE-ISSUED: October 21, 1997

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Flen.o slashed. ; Bent Stenl.o slashed.se		N/A	N/A	DKX
Christensen; Ib	Aller.o slashed.d	N/A	N/A	DKX
Larsen; Robert	Virum	N/A	N/A	DKX
Johansen; Steffen	Ega	N/A	N/A	DKX
Radich				
Johnson; Eric A.	Madison	WI	N/A	N/A

US-CL-CURRENT: 435/255.1; 435/67

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KVMC	Draw. Desc	Image
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☐ 9. Document ID: US 5670548 A

L13: Entry 9 of 19

File: USPT

Sep 23, 1997

US-PAT-NO: 5670548

DOCUMENT-IDENTIFIER: US 5670548 A

TITLE: Pigmentation with carotenoids

DATE-ISSUED: September 23, 1997

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Bernhard; Kurt	Lupsingen	N/A	N/A	CHX
Broz; Jiri	Rheinfelden	N/A	N/A	CHX
Hengartner; Urs	Basel	N/A	N/A	CHX
Kreienbuhl; Paul	Riehen	N/A	N/A	CHX
Schiedt; Katharina	Arlesheim	N/A	N/A	CHX

US-CL-CURRENT: 514/725; 424/442

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 10. Document ID: US 5605699 A

L13: Entry 10 of 19

File: USPT

Feb 25, 1997

US-PAT-NO: 5605699

DOCUMENT-IDENTIFIER: US 5605699 A

TITLE: Pigmentation with carotenoids

DATE-ISSUED: February 25, 1997

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Bernhard; Kurt	Lupsingen	N/A	N/A	CHX
Broz; Jiri	Rheinfelden	N/A	N/A	CHX
Hengartner; Urs	Basel	N/A	N/A	CHX
Kreienbuhl; Paul	Riehen	N/A	N/A	CHX
Schiedt; Katharina	Arlesheim	N/A	N/A	CHX

US-CL-CURRENT: 424/442; 424/489, 426/807

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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Terms	Documents
(crustacean? and fish)and (feedstuff)	19

Documents, starting with Document:

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**WEST**[Generate Collection](#)**Search Results - Record(s) 11 through 19 of 19 returned.**☐ 11. Document ID: US 5599711 A

L13: Entry 11 of 19

File: USPT

Feb 4, 1997

US-PAT-NO: 5599711

DOCUMENT-IDENTIFIER: US 5599711 A

TITLE: Astaxanthin-producing yeast cells, methods for their preparation and their use

DATE-ISSUED: February 4, 1997

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP	CODE	COUNTRY
Flen.o slashed.; Bent	Stenl.o slashed.se	N/A	N/A		DKX
Christensen; Ib	Aller.o slashed.d	N/A	N/A		DKX
Larsen; Robert	Virum	N/A	N/A		DKX
Johansen; Steffen R.	Ega	N/A	N/A		DKX
Johnson; Eric A.	Madison	WI	N/A		N/A

US-CL-CURRENT: 435/255.1; 435/67

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 12. Document ID: US 5356810 A

L13: Entry 12 of 19

File: USPT

Oct 18, 1994

US-PAT-NO: 5356810  
DOCUMENT-IDENTIFIER: US 5356810 A

TITLE: Astaxanthin-producing yeast cells, methods for their preparation and their use

DATE-ISSUED: October 18, 1994

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Fleno; Bent	Stenlose	N/A	N/A	DKX
Christensen; Ib	Allerod	N/A	N/A	DKX
Larsen; Robert	Virum	N/A	N/A	DKX
Johansen; Steffen R.	Ega	N/A	N/A	DKX
Johnson; Eric A.	Madison	WI	N/A	N/A

US-CL-CURRENT: 435/255.1; 435/67

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 13. Document ID: US 5356809 A

L13: Entry 13 of 19 File: USPT Oct 18, 1994

US-PAT-NO: 5356809  
DOCUMENT-IDENTIFIER: US 5356809 A

TITLE: Processes for in vivo production of astaxanthin and phaffia rhodozyma yeast of enhanced astaxanthin content

DATE-ISSUED: October 18, 1994

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Johnson; Eric A.	Madison	WI	N/A	N/A
Yang; Huei-hsiung	Rockville	MD	N/A	N/A
Geldiay-Tuncer; Beril	Bostanli-Karsiyaka	N/A	N/A	TRX
Hall; William T.	Rockville	MD	N/A	N/A
Schreiber; David	Columbia	MD	N/A	N/A
Ho; Kwok	San Diego	CA	N/A	N/A

US-CL-CURRENT: 435/255.1; 435/244, 435/256.8, 435/67, 435/911

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 14. Document ID: US 5182208 A

L13: Entry 14 of 19 File: USPT Jan 26, 1993

US-PAT-NO: 5182208  
DOCUMENT-IDENTIFIER: US 5182208 A

TITLE: Processes for in vivo production of astaxanthin and  
phaffia rhodozyma yeast of enhanced astaxanthin content

DATE-ISSUED: January 26, 1993

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Johnson; Eric A.	Madison	WI	N/A	N/A
Schreiber; David	Columbia	MD	N/A	N/A
Ho; Kwok P.	Columbia	MD	N/A	N/A
Hall; William T.	Rockville	MD	N/A	N/A
Yang; Huei-hsiung	Rockville	MD	N/A	N/A
Geldiay-Tuncer; Beril	College Park	MD	N/A	N/A

US-CL-CURRENT: 435/255.1; 435/244, 435/253.6, 435/67, 435/911

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 15. Document ID: US 5030657 A

L13: Entry 15 of 19 File: USPT Jul 9, 1991

US-PAT-NO: 5030657  
DOCUMENT-IDENTIFIER: US 5030657 A

TITLE: L-carnitine supplemented catfish diet

DATE-ISSUED: July 9, 1991

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Burtle; Gary J.	Tifton	GA	N/A	N/A
Newton; G. Larry	Tifton	GA	N/A	N/A
Blum; Stephen A.	Des Moines	IA	N/A	N/A

US-CL-CURRENT: 514/556; 426/805, 514/951, 514/963

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 16. Document ID: US 4958011 A

L13: Entry 16 of 19 File: USPT Sep 18, 1990

US-PAT-NO: 4958011  
DOCUMENT-IDENTIFIER: US 4958011 A

TITLE: Ester-stabilized chitin

DATE-ISSUED: September 18, 1990

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Bade; Maria L.	Lexington	MA	02173	N/A

US-CL-CURRENT: 536/20

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 17. Document ID: US 4906479 A

L13: Entry 17 of 19 File: USPT Mar 6, 1990

US-PAT-NO: 4906479  
DOCUMENT-IDENTIFIER: US 4906479 A

TITLE: Feedstuff for artemia

DATE-ISSUED: March 6, 1990

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Kitagawa; Kiyohiro	Chiba	N/A	N/A	JPX
Kojima; Eiji	Kanagawa	N/A	N/A	JPX
Seto; Akira	Kanagawa	N/A	N/A	JPX
Sakamoto; Shuichi	Tokyo	N/A	N/A	JPX
Nozawa; Takuji	Chiba	N/A	N/A	JPX

US-CL-CURRENT: 426/1; 426/2, 426/59, 426/60

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 18. Document ID: US 4634592 A

L13: Entry 18 of 19 File: USPT Jan 6, 1987

US-PAT-NO: 4634592  
DOCUMENT-IDENTIFIER: US 4634592 A

TITLE: Process for the manufacture of aquatic bait blocks

DATE-ISSUED: January 6, 1987

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Faber; Jerry L.	Decatur	IN	N/A	N/A
Kent, Jr.; Richard W.	Ft. Wayne	IN	N/A	N/A

US-CL-CURRENT: 426/1; 426/658, 426/805

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 19. Document ID: US 3989847 A

L13: Entry 19 of 19

File: USPT

Nov 2, 1976

US-PAT-NO: 3989847

DOCUMENT-IDENTIFIER: US 3989847 A

TITLE: Process for preparation of seasonings derived from animal meat

DATE-ISSUED: November 2, 1976

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Kurihara; Shinji	Fukuyama, Hiroshima	N/A	N/A	JA
Osazima; Kazuharu	Fukuyama, Hiroshima	N/A	N/A	JA

US-CL-CURRENT: 426/7; 426/533, 426/55

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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Terms	Documents
(crustacean? and fish)and (feedstuff)	19

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☐ 1. Document ID: US 6184027 B1

US-CL-CURRENT: 435/261

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KMNC	Draw. Desc	Image
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☐ 2. Document ID: US 6159719 A

US-CL-CURRENT: 435/206; 435/18, 435/7.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KMC	Draw Desc	Image
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☐ 3. Document ID: US 6156568 A

L14: Entry 3 of 8

File: USPT

Dec 5, 2000

US-PAT-NO: 6156568

DOCUMENT-IDENTIFIER: US 6156568 A

TITLE: Transformed eukaryotic cells

DATE-ISSUED: December 5, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Cooper; Richard K.	Baton Rouge	LA	N/A	N/A
Enright; Frederick M.	Baton Rouge	LA	N/A	N/A

US-CL-CURRENT: 435/325; 435/320.1, 435/410, 435/455, 435/468,  
435/6, 435/69.1, 536/23.1, 536/24.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☒ 4. Document ID: US 6143883 A

L14: Entry 4 of 8

File: USPT

Nov 7, 2000

US-PAT-NO: 6143883

DOCUMENT-IDENTIFIER: US 6143883 A

TITLE: Water-soluble low molecular weight beta-glucans for  
modulating immunological responses in mammalian system

DATE-ISSUED: November 7, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Lehmann; Joachim	Scottsdale	AZ	N/A	N/A
Kunze; Rudolf	Berlin	N/A	N/A	DEX

US-CL-CURRENT: 536/55.3; 435/18, 435/7.31, 536/123.1, 536/123.12,  
536/124, 536/127, 536/55.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 5. Document ID: US 6090573 A

L14: Entry 5 of 8

File: USPT

Jul 18, 2000

US-PAT-NO: 6090573

DOCUMENT-IDENTIFIER: US 6090573 A

TITLE: Detecting eubacteria and fungus and determining their antibiotic sensitivity by using catalytically inactive murein binding enzymes

DATE-ISSUED: July 18, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Laine; Roger A.	Baton Rouge	LA	70808	N/A
Lo; Wai Chun Jennifer	Baton Rouge	LA	70808	N/A

US-CL-CURRENT: 435/32; 435/18, 435/206, 435/29, 435/34

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 6. Document ID: US 5998698 A

L14: Entry 6 of 8

File: USPT

Dec 7, 1999

US-PAT-NO: 5998698

DOCUMENT-IDENTIFIER: US 5998698 A

TITLE: Transgenic fish capable of expressing exogenous lytic peptides

DATE-ISSUED: December 7, 1999

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Cooper; Richard K.	Baton Rouge	LA	N/A	N/A
Enright; Frederick M.	Baton Rouge	LA	N/A	N/A

US-CL-CURRENT: 800/20

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 7. Document ID: US 5935804 A

L14: Entry 7 of 8

File: USPT

Aug 10, 1999



US-PAT-NO: 5935804

DOCUMENT-IDENTIFIER: US 5935804 A

TITLE: Method for detecting eubacteria in biological samples with catalytically inactive murein binding enzymes

DATE-ISSUED: August 10, 1999

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Laine; Roger A.	Baton Rouge	LA	70808	N/A
Lo; Wai C. J.	Baton Rouge	LA	70808	N/A

US-CL-CURRENT: 435/18; 435/206, 435/29

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 8. Document ID: US 5719055 A

L14: Entry 8 of 8

File: USPT

Feb 17, 1998

US-PAT-NO: 5719055

DOCUMENT-IDENTIFIER: US 5719055 A

TITLE: Transposon-based transformation vectors

DATE-ISSUED: February 17, 1998

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Cooper; Richard K.	Baton Rouge	LA	N/A	N/A

US-CL-CURRENT: 435/320.1; 435/252.33, 536/23.2, 536/23.7,  
536/24.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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Terms	Documents
(crustacean? and fish)and (lipopolysaccharide)	8

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Documents, starting with Document:

[8](#)**Display Format:**[CIT](#)[Change Format](#)

**WEST**[Generate Collection](#)**Search Results - Record(s) 1 through 10 of 65 returned.**☐ 1. Document ID: US 6210567 B1

L2: Entry 1 of 65

File: USPT

Apr 3, 2001

US-PAT-NO: 6210567

DOCUMENT-IDENTIFIER: US 6210567 B1

TITLE: Filtration device for tank water for aquarium fish

DATE-ISSUED: April 3, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Takada; Shunsuke	Meguro-ku, Tokyo	N/A	N/A	JPX

US-CL-CURRENT: 210/169; 119/227, 119/260, 210/252, 210/282,  
210/287, 210/336

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 2. Document ID: US 6160023 A

L2: Entry 2 of 65

File: USPT

Dec 12, 2000

US-PAT-NO: 6160023

DOCUMENT-IDENTIFIER: US 6160023 A

TITLE: Use of bronopol for the treatment of diseases in fish

DATE-ISSUED: December 12, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Braidwood; Julian Charles	Cumbria	N/A	N/A	GBX

US-CL-CURRENT: 514/727

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 3. Document ID: US 6156568 A

L2: Entry 3 of 65

File: USPT

Dec 5, 2000

US-PAT-NO: 6156568

DOCUMENT-IDENTIFIER: US 6156568 A

TITLE: Transformed eukaryotic cells

DATE-ISSUED: December 5, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Cooper; Richard K.	Baton Rouge	LA	N/A	N/A
Enright; Frederick M.	Baton Rouge	LA	N/A	N/A

US-CL-CURRENT: 435/325; 435/320.1, 435/410, 435/455, 435/468,  
435/6, 435/69.1, 536/23.1, 536/24.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 4. Document ID: US 6139731 A

L2: Entry 4 of 65

File: USPT

Oct 31, 2000

US-PAT-NO: 6139731

DOCUMENT-IDENTIFIER: US 6139731 A

TITLE: Iodinated water treatment process

DATE-ISSUED: October 31, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Harvey; Wayne A.	Dartmouth	N/A	N/A	CAX
Mullins; Terence F.	Dartmouth	N/A	N/A	CAX
MacDonald; Daniel J.	Dartmouth	N/A	N/A	CAX

US-CL-CURRENT: 210/175; 210/192, 210/205, 210/742

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 5. Document ID: US 6086755 A

L2: Entry 5 of 65

File: USPT

Jul 11, 2000

US-PAT-NO: 6086755

DOCUMENT-IDENTIFIER: US 6086755 A

TITLE: Floating hydroponic biofiltration device

DATE-ISSUED: July 11, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Tepper; Julius	Manoville	NY	11949	N/A

US-CL-CURRENT: 210/150; 119/260, 210/169, 210/170, 210/242.1,  
210/602, 47/64

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 6. Document ID: US 6065430 A

L2: Entry 6 of 65

File: USPT

May 23, 2000

US-PAT-NO: 6065430

DOCUMENT-IDENTIFIER: US 6065430 A

TITLE: Fish culturing system

DATE-ISSUED: May 23, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Sheriff; Richard L.	Chester	PA	21619	N/A

US-CL-CURRENT: 119/227

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw Desc	Image
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☐ 7. Document ID: US 6062170 A

L2: Entry 7 of 65

File: USPT

May 16, 2000

US-PAT-NO: 6062170  
DOCUMENT-IDENTIFIER: US 6062170 A

TITLE: Method and apparatus for plankton reduction in fish farming

DATE-ISSUED: May 16, 2000

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Finch; Richard John	Campbell River	N/A	N/A	CAX
Davidson; D'Arcy	Victoria	N/A	N/A	CAX
Lueck; Rolf	Victoria	N/A	N/A	CAX

US-CL-CURRENT: 119/215; 114/222, 119/240

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 8. Document ID: US 6041738 A

L2: Entry 8 of 65 File: USPT Mar 28, 2000

US-PAT-NO: 6041738  
DOCUMENT-IDENTIFIER: US 6041738 A

TITLE: Fish pond methods and systems

DATE-ISSUED: March 28, 2000

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Hemauer; Thomas Joseph	Chilton	WI	N/A	N/A
Niquette; Daniel John	Chilton	WI	N/A	N/A

US-CL-CURRENT: 119/226; 119/227

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☐ 9. Document ID: US 6025008 A

L2: Entry 9 of 65 File: USPT Feb 15, 2000

US-PAT-NO: 6025008  
DOCUMENT-IDENTIFIER: US 6025008 A

TITLE: Yogurt

DATE-ISSUED: February 15, 2000

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Akahoshi; Ryoichi	Tokyo	N/A	N/A	JPX
Mizobuchi; Takahiro	Tokyo	N/A	N/A	JPX
Takahashi; Yoshihiro	Tokyo	N/A	N/A	JPX
Saita; Taketsugu	Tokyo	N/A	N/A	JPX

US-CL-CURRENT: 426/583; 426/34, 426/580, 426/585

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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☒ 10. Document ID: US 6017722 A

L2: Entry 10 of 65

File: USPT

Jan 25, 2000

US-PAT-NO: 6017722

DOCUMENT-IDENTIFIER: US 6017722 A

TITLE: Luminous bacteria and methods for the isolation,  
identification and quantitation of toxicants

DATE-ISSUED: January 25, 2000

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Becvar; James E.	El Paso	TX	N/A	N/A
Becvar; Laura E.	El Paso	TX	N/A	N/A

US-CL-CURRENT: 435/8; 210/658, 422/52, 422/61, 422/68.1, 435/29,  
435/909, 436/162, 436/172

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw Desc	Image
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Generate Collection

Terms	Documents
(gram negative bacteria)[AB] AND (fish)[AB]	65

Display

10

Documents, starting with Document: 11

**WEST****End of Result Set**

Generate Collection

L22: Entry 1 of 1

File: USPT

Sep 17, 1996

US-PAT-NO: 5556625DOCUMENT-IDENTIFIER: US 5556625 A

TITLE: Pharmaceutical composition comprising Rosa roxburghii, Artemisi argyi folium and Brassica oleracea var. capitata L. used to reduce the symptoms of diarrhea

DATE-ISSUED: September 17, 1996

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Araki; Seiichi	Ibaraki	N/A	N/A	JPX
Suzuki; Mamoru	Ibaraki	N/A	N/A	JPX
Fujimoto; Masatoshi	Ibaraki	N/A	N/A	JPX
Ueki; Tadashi	Tokyo	N/A	N/A	JPX

## ASSIGNEE-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY	TYPE CODE
Eisai Co., Ltd.	Tokyo	N/A	N/A	JPX	03

APPL-NO: 8/ 433312

DATE FILED: May 3, 1995

## PARENT-CASE:

This application is a continuation of U.S. Ser. No. 08/133,708, filed 7, 1993, now abandoned.

## FOREIGN-APPL-PRIORITY-DATA:

COUNTRY	APPL-NO	APPL-DATE
JP	4-291995	October 7, 1992
JP	4-291996	October 7, 1992
JP	5-144345	May 25, 1993
JP	5-206808	July 30, 1993

INT-CL: [6] A61K 35/78

US-CL-ISSUED: 424/195.1; 424/93.7, 514/867

US-CL-CURRENT: 424/740; 424/93.7, 514/867

FIELD-OF-SEARCH: 424/93.7, 424/195.1, 514/867

PRIOR-ART-DISCLOSED:

## U.S. PATENT DOCUMENTS

☐ Search Selected☐ Search ALL

PAT-NO	ISSUE-DATE	PATENTEE-NAME	US-CL
<input type="checkbox"/> 3108040	October 1963	Folkers	167/55
<input type="checkbox"/> 5091195	February 1992	Havens	426/2
<input type="checkbox"/> 5162037	November 1992	Whitson-Fischman	600/12

## OTHER PUBLICATIONS

Shabana, M. M. et al., "Egypt. J. Pharm.," vol. 19, (#1-4), pp. 271-28 1978.  
Ishida, H et al. "Yakugaku Zasshi," vol. 109 (#3), pp. 179-183, 1989 (translation).  
Yoshida, T., et al., "Chem. Pharm. Bull.," vol. 35, #5, 1987, pp. 1817-1822.  
Lao, A, et al., "Chem. Pharm. Bull.," vol. 32, #2, 1984, pp. 723-727.  
Agris International, Abstract No. 89-052859, Chengyuan et al: "A Study the Value of Cili (Rosa roxburghii Tratt) Residue as an Animal Feed" & Journal of Guizhou Agricultural Science Jun. 1988, vol. 3, pp. 45-48.

ART-UNIT: 188

PRIMARY-EXAMINER: Wityshyn; Michael G.

ASSISTANT-EXAMINER: Larson; Kristin

ATTY-AGENT-FIRM: Flynn, Thiel, Boutell &amp; Tanis, P.C.

## ABSTRACT:

A method of reducing the symptoms of diarrhea by administering to an animal a composition containing Rosa roxburghii, Artemisiae argyi foli and Brassica oleracea var. capitata L.

4 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

## BRIEF SUMMARY:

## BACKGROUND OF THE INVENTION

## 1. Field of the Invention

The present invention relates to a medicinal agent or food which potentiates the immune system of an animal, including a human being, thereby protect the animal from infection, and a feed or feedstuff hav an immunopotentiating activity.

The present invention also relates to a medicinal agent or food for regulating a function of the digestive tract or gastrointestinal of an



animal, including a human being, and a feed or feedstuff having the activity of regulating the function of the digestive tract or gastrointestinal of an animal.

Further, the present invention relates to a medicinal agent for improv antibiotic absorption by an animal, including a human being, and a fee feedstuff for improving antibiotic absorption by an animal.

Furthermore, the present invention relates to a medicinal agent for accelerating an animal growth or for improving an egg production rate, weight, egg quality or eggshell strength of an animal and a feed or feedstuff having the activity of accelerating an animal growth or the activity of improving an egg production rate, egg weight, egg quality eggshell strength of an animal.

## 2. Description of the Related Art

With the recent progress in immunology, it has come to be thought that various maladies and infectious diseases of an animal including a huma being, are caused by a weakening or deficiency in the immune system of animal.

For example, a human being frequently suffers from a weakening or deficiency in his immune system because of bronchial asthma, allergic diseases, articular rheumatism, autoimmune diseases, nutritional disorders, surgical operation, aging, cancer, organ transplantation, pregnancy or the like, resulting in the complication of an infectious disease such as respiratory infections, sepsis or urinary infections.

Up to this time, various antibiotics have been administered to patient with such maladies or infectious diseases. Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandr and aquaculture in order to raise livestock, poultries or fish efficiently, and in such raising, it has been also a practice to administer a high dose of an antibiotic.

The repeated administration of an antibiotic for a long period causes generation of antibiotic resistant bacteria to lower the effect of the antibiotic. Further, hospital infection has also become a significant problem. Under these circumstances, it has been expected to develop a preventive and therapeutic agent which can potentiate the immune syste while reducing the dosage of an antibiotic.

Further the overcrowded raising employed in the fields of animal husba and aquaculture has a problem in that various infectious diseases frequently break out because of stress or Juvenile immuno-deficiency. Furthermore, when a high dose of an antibiotic is administered as a countermeasure against the problem, there occur other problems, for example the antibiotic is not completely consumed and antibiotic resis bacteria propagate in the environment.

Recently, animals, including human beings, have frequently suffered fr various gastrointestinal diseases which are caused by a lowering or deficiency in the immune system, stress, dyspepsia or the like, and mo of which are accompanied with diarrhea.

For example, a human being becomes susceptible to an infectious diseas

the digestive tract as his resistance lowers, and representative examples of the infectious diseases include bacterial, viral and parasitic diarrhea. Further, the above gastrointestinal diseases also include acute diarrhea caused by food poisoning and food allergy and chronic diarrhea caused by disorder of digestion and absorption, excess gut hormone and colic diseases.

It has been the practice to administer an intestinal depressomotor for intestines, an astringent, an irritant-absorbing agent, a torpent for enteric mucous membranes or various antibiotics against these gastrointestinal diseases.

Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandry and aquaculture to raise livestock, poultry and fish efficiently. In such raising, it has been a practice to administer the above therapeutic agents for the treatment of diarrhea caused by stress or dyspepsia. Particularly, a large amount of an antibiotic has been used for the prevention of infectious diseases.

The administration of the above depressomotor or astringent is essentially a nosotropic means, while that of an antibiotic is an etiotropic one. However, the repeated administration of an antibiotic for the prevention of infectious diseases has a problem that the preventive effect lowers owing to the generation of antibiotic resistant bacteria.

Under these circumstances, it has been expected to develop a novel preventive and therapeutic agent which can regulate gastrointestinal functions themselves and is safe.

Up to this time, various antibiotics have been administered to patients with various maladies or infectious diseases. Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandry and aquaculture to raise livestock, poultry and fish efficiently. In such raising, it has been a practice to administer a high dose of an antibiotic.

With respect to the use of an antibiotic, however, there have been problems such that the antibiotic accumulates in the body of an animal to affect the body and that the administration of a more potent antibiotic becomes necessary because of the generation of antibiotic resistant bacteria. Further, the administration of an antibiotic to an animal is problematic also in that the environment is polluted with the antibiotic contained in the excrement of the animal. Under these circumstances, it has been expected that the dose of the antibiotic to be administered is reduced. In order to reduce the dose of the antibiotic, it has been believed effective that the absorption of the antibiotic through the digestive tract should be enhanced to attain an effect equal or superior to that attained by the administration thereof in a high dose, even when the antibiotic is administered in a low dose. However, no substance which can improve the absorption of an antibiotic has been found as yet.

In preparing animal feed, various attempts have been widely made at improving the feeding efficiency for animals and also at accelerating growth thereof by adding various antibiotics to the feed, incorporating increased amounts of proteins into the feed, changing the feeding method, improving the dosage form of the feed.

With respect to the use of an antibiotic, however, there have been pointed out problems such that the antibiotic remains or accumulates in the body of an animal to affect the body and that the administration of a more potent antibiotic becomes necessary because of the generation of resistant bacteria. Further, the administration of an antibiotic to an animal is problematic also in that the environment is polluted with the antibiotic contained in the excrement of the animal. In addition, the effects obtained by varying the feeding method and the dosage form of the feed are limited. Under these circumstances, it is necessary to develop a safe and less problematic animal growth accelerator, feed and improver for the production ratio, egg weight, egg quality or eggshell strength of animals.

## DISCLOSURE OF THE INVENTION

### Summary of the Invention

In view of the above problems, the present inventors have extensively studied for many years and have found that *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. are useful for overcoming the problems. The present invention has been accomplished on the basis of this finding.

Thus, the present invention provides;

(1) a pharmaceutical composition comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(2) a pharmaceutical composition comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.;

(3) a feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.;

(4) a feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a basal diet;

(5) a method for feeding a feedstuff comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to an animal: and

(6) a method for feeding a feedstuff comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a basal diet to an animal.

In view of the above problems with respect to the use of antibiotics, the present inventors have extensively studied for many years on protective agents which are safe for animals and have found that *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. have an immunopotentiating activity. The present invention has been accomplished on the basis of this finding.

Thus, the present invention provides;

(7) a method for immunopotentiating and protecting from infectious diseases, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* to an animal;

(8) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for preparing a medicine for an animal for immunopotentiating and protecting from infectious diseases;

(9) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for immunopotentiating and protecting from infectious diseases in an animal;

(10) a pharmaceutical composition for immunopotentiating and protectin from infectious diseases in an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* and a pharmaceutically acceptable carrier;

(11) a medicinal agent for immunopotentiating and protecting from infectious diseases of an animal, comprising a substance selected from group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brass oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier; a

(12) a feedstuff for an animal, which is useful for immunopotentiating protecting from infectious diseases in an animal, comprising a substan selected from the group consisting of *Rosa roxburghii*, *Artemisiae argy folium* and *Brassica oleracea* var. *capitata* L.

In view of the above problem, the present inventors have extensively studied for many years regulating agents for the digestive tract which safe for human beings and animals, and have found that *Rosa roxburghii* *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. have th activity of regulating gastrointestinal functions. The present inventi has been accomplished on the basis of this finding.

Thus, the present invention provides;

(13) a method for regulating the digestive tract, which comprises administering a pharmacologically effective amount of a substance sele from the group consisting of *Rosa roxburghii*, *Artemisiae argyl folium* *Brassica oleracea* var. *capitata* L. to an animal;

(14) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for preparing a medicine for an animal for regulating the digestive tr

(15) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for regulating the digestive tract of an animal;

(16) a pharmaceutical composition for regulating the digestive tract of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(17) a medicinal agent for regulating the digestive tract of an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* and a pharmaceutically acceptable carrier; and

(18) a feedstuff for an animal, which is useful for regulating the digestive tract of the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L.

To provide an absorbefacient which enables a reduction in the dose of antibiotic when administered together with the antibiotic, the present inventors have extensively studied to find out that the absorption of antibiotic can surprisingly be improved when *Rosa roxburghii* alone or or more members selected from among *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. are administered simultaneously with the antibiotic or before or after the administration of the antibiotic.

Thus, the present invention provides;

(19) a method for improving antibiotic absorption, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. to an animal;

(20) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* for preparing a medicine for an animal for improving its antibiotic absorption;

(21) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* for improving antibiotic absorption in an animal;

(22) a pharmaceutical composition for improving antibiotic absorption in an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(23) a medicinal agent for improving antibiotic absorption in an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* and a pharmaceutically acceptable carrier; and

(24) a feedstuff for an animal, which is useful for improving antibiotic absorption in the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L.

To provide a feedstuff free from the abovementioned defects of an ordinary feedstuff and a method of administering the feedstuff to animals, the present inventors have extensively studied. As a result, they have found that the growth of animals, including fetuses, is surprisingly accelerated by administering *Rosa roxburghii* alone or two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to them. The inventors have further found that an improvement in the egg production rate, egg weight, egg quality or eggshell strength of birds and fishes is also accelerated.

Thus, the present invention provides;

(25) a method for accelerating growth of an animal, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to the animal;

(26) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. for preparing a medicine for an animal for accelerating growth of the animal;

(27) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. for accelerating growth of an animal;

(28) a pharmaceutical composition for accelerating the growth of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(29) a medicinal agent for accelerating the growth of an animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(30) a feedstuff for an animal, which is useful for accelerating the growth of the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.;

(31) a method for improving an egg production rate, egg weight or eggshell strength of an animal, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to the animal;

(32) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. for preparing a medicine of an animal for improving an egg production rate, egg weight, egg quality or eggshell strength of the animal;

(33) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. for improving an egg production rate, egg weight, egg quality or eggshell strength of the animal;

strength of an animal;

(34) a pharmaceutical composition for improving an egg production rate egg weight, egg quality or eggshell strength of an animal comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. and a pharmaceutically acceptable carrier;

(35) a medicinal agent for improving an egg production rate, egg weight egg quality or eggshell strength of an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. and a pharmaceutically acceptable carrier; and

(36) a feedstuff for an animal, which is useful for improving an egg production rate, egg weight, egg quality or eggshell strength of the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata*

Further scope and applicability of the present invention will become apparent from the detailed description given hereinafter. However, it should be understood that the detailed description and specific examples while indicating preferred embodiments of the invention, are given by way of illustration only, since various changes and modifications within the spirit and scope of the invention will become apparent to those skilled in the art from this detailed description.

#### DETAILED DESCRIPTION OF THE INVENTION

*Rosa roxburghii* is a perennial shrub of the family Rosaceae and native to Guizhou in China and its fruit has been noticed as the material of jui jam or liquor. The fruit of *Rosa roxburghii* has been known to have a pharmacological activity and is useful as an antiulcer agent by virtue of its cancer-preventing, cholesterol level-lowering and antistress activities.

*Artemisiae argyi folium* is a plant of the family Compositae and has been known to be useful as an antidiarrheal or antiabdominalgia agent, a hemostatic or the like. Further, this plant has been known to exhibit low antimicrobial activity only against Gram-positive bacteria. *Artemisiae argyi folium* includes, for example, *Artemisia princeps* Pampanini, *Artemisia mongolica* Fischer, *Artemisia argyi* LEVL. et VANT., and *Artemisia lavandulaefolia* DC.

*Brassica oleracea var. capitata* L. is a plant of the family Cruciferae and has been used as food.

The *Rosa roxburghii* to be used in the present invention is not particularly limited in form, but may have any form so far as it contains the essences of *Rosa roxburghii*. Generally, the fruit of *Rosa roxburghii* may be used in its raw state or as a dry powder prepared by conventional means prior to use, or an extract prepared by using water, an organic solvent or a mixture of both as an extractant.

The *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. to be used in the present invention are not particularly limited in form, but

may have any form so far as it contains the essences thereof. Generally the leaf of *Artemisiae argyi folium* or *Brassica oleracea var. capitata* may be used in a raw state or as a dry powder prepared by a conventional means prior to use, or an extract prepared by using water, an organic solvent or a mixture of both as an extractant.

An extract from the above leaves can be prepared by, e.g., a process which comprises immersing 1 part by weight of the raw material in 5 parts by weight of water, boiling the obtained mixture under heating for 30 min to conduct extraction, filtering the resulting system, and concentrating the obtained filtrate to 3.6 parts by weight. The obtained extract may be powdered by spray drying, freeze drying, vacuum drying (vacuum concentration) or the like.

When an organic solvent is used for extraction, methanol, ethanol, n-propanol, n-butanol, acetone, ethyl acetate, ether, methylene chloride, chloroform, benzene, carbon tetrachloride and petroleum ether are preferable. These organic solvents may be used alone or as a mixture of two or more of them.

The extracts thus produced may be used as such or may be concentrated, diluted or freed from the solvent prior to use.

The extract of *Rosa roxburghii* to be used in the present invention may be one commercially available under the trade name of "Rosa roxburghii extract powder MF"; which is a product of Maruzen Seiyaku K. K. comprising 30% of an extract of *Rosa roxburghii* and 70% of dextrin.

In the present invention, at least one of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. is used.

When *Rosa roxburghii* and *Artemisiae argyi folium* are used in the present invention, the ratio between them is not particularly limited. Generally *Artemisiae argyi folium* is used, in terms of raw leaf, that is, when prescribed as the weight of raw leaf, in an amount of 0.25 to 400 parts by weight, preferably 0.5 to 200 parts by weight, still preferably 1 to 10 parts by weight, based on 1 part by weight of the extract of *Rosa roxburghii*.

When both *Artemisiae argyi folium* and *Brassica oleracea var. capitata* are used together with *Rosa roxburghii*, the amounts thereof are not particularly limited. Generally, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. are used in amounts of 0.25 to 400 parts by weight and 0.5 to 800 parts by weight, respectively, in terms of their respective raw leaf, based on 1 part by weight of the extract of *Rosa roxburghii*. It is preferable that *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. be used in amounts of 0.5 to 200 parts by weight and 1 to 400 parts by weight respectively, still preferably in amounts of 1 to 100 parts by weight and 5 to 200 parts by weight respectively.

The pharmaceutical composition or the medicinal agent of the present invention may be administered, with the purpose of the prevention of diseases, as a food having a regulating effect on a living body, i.e., so-called functional food, which can be prepared by adding this composition or agent to food.

When the pharmaceutical composition or the medicinal agent of the present



invention comprising one or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* is used as a medicine or so-called health food, it may be in the form tablet, granule, powder, capsule or syrup. The pharmaceutical composition or the medicinal agent is prepared by mixing the members with a conventional filler or carrier, binder, lubricant or the like and treating the obtained mixture in the conventional manner.

When the pharmaceutical composition or the medicinal agent of the present invention comprising one or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* is used for an animal, such as a mammal, a bird and a fish, especially a livestock, the dosage form thereof is not particularly limited. For example, it may be administered to livestock in a state mixed with a basal diet (or feed or feedstuff). That is, this composition or agent may be mixed with a basal diet (or feed or feedstuff) just before using or may be premixed with a basal diet (or feed or feedstuff). In other words, the feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. of the present invention may be administered to an animal as a feed having a biophylactic and regulating function.

The arbitrary basal diet for animals, which is used for preparing the feedstuff according to the present invention, is not particularly limited. Examples of the raw materials constituting the basal diet include grains such as corn, milo and wheat flour, brans such as defatted rice bran and wheat bran, animal substances such as fish meal and skim milk, vegetable oil cake such as soybean oil cake, and additives such as calcium carbonate, calcium phosphate, common salt, vitamin B<sub>12</sub>, DL-methionine, choline chloride, manganese sulfate, dry iron sulfate, calcium iodate, copper sulfate, dry zinc sulfate and sodium saccharin. The basal diet may be prepared by blending some members selected from among these materials. The formulation of the basal diet varies depending upon the animal to which the diet is administered.

The improved feed of the present invention, i.e., the feedstuff containing at least one of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. can be administered to various livestock, poultry, pets and fish. Livestock includes pig, cattle, horse, goat, sheep and rabbit; poultry includes chicken, Japanese quail, turkey and duck; pets includes dogs and cats; and fish includes yellowtail, sea bream, flatfish, globefish, hardtail, amberjack, salmon, carp, eel, sweetfish, trout, char and lobster.

Needless to say, the pharmaceutical composition, the medicinal agent and the feedstuff of the present invention are non-toxic.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for immunopotentiating and protecting from infectious diseases.

In this case, *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. may be used alone or as a combination of two or more of them. In other words, the combination of *Rosa roxburghii* with *Artemisiae argyi folium*, that of *Rosa roxburghii* with *Brassica oleracea* var. *capitata* L., that of *Artemisiae argyi folium* with *Brassica oleracea* var. *capitata* L., and that of *Rosa roxburghii* with *Artemisiae argyi fo*

and *Brassica oleracea* var. *capitata* L. can be used.

*Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. exhibit an immunopotentiating and protective activity even when used each alone. However, the use of a combination of two or more of them surprisingly exhibits an immunopotentiating and protective activity which is higher than that of the use of each of them.

The pharmaceutical composition or the medicinal agent of the present invention can safely and effectively potentiate the immune function of livestock such as cattle, pigs or horses, poultry such as chicken or Japanese quail, fish such as young yellowtail, sea bream, eel, trout, or goldfish, pets such as dogs or cats, or a human being to protect the animal or the like -From various infectious diseases, which is one of the objects of the present invention.

The term "immunopotentiation" used in this specification means "potentiation of the immune function of an animal such as a human being, mammal, fish or the like".

The pharmaceutical composition or the medicinal agent of the present invention have the effect of enhancing the immune function of an animal including a human being, and serves as a preventive and therapeutic pharmaceutical composition or agent for various maladies and infectious diseases by virtue of this effect, so that the diseases against which the composition or agent is efficacious are not particularly limited. For example, it is efficacious against, e.g., articular rheumatism, autoimmune diseases, bronchial asthma, nutritional disorders, surgical operation, diseases of old age and various infectious diseases such as respiratory infection, sepsis and urinary infection with respect to a human being.

With respect to animals other than the humans, this composition or agent is efficacious against scours, epizootic pneumonia, atrophic rhinitis, infectious enterogastroenteritis of a pig, pneumonia and Marek's disease of chicken, scours, pneumonia and mastitis of cattle, and AIDS and leukemia of a pet.

Further, with respect to fish, the infectious disease against which the composition or agent of the present invention is efficacious is not particularly limited and includes bacterial diseases such as streptococcal and nodosity, and viral diseases.

In this case, the dose of *Rosa roxburghii* to be administered varies depending upon the dosage form and the subject animal, so that it is not particularly limited.

For example, an extract of *Rosa roxburghii* is administered to livestock such as a pig in a dose of 25 mg or above, preferably 50 mg or above, still preferably 100 mg or above per kilogram of the body weight.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for regulating a function of the digestive tract.

In this case, *Rosa roxburghii* is preferably used as the essential component. The activity of regulating the function(s) of the digestive tract, i.e., the gastrointestinal function(s) regulating activity, of

pharmaceutical composition or the medicinal agent is synergistically enhanced when the composition or agent further contains *Artemisiae arg folium* or *Artemisiae argyl folium* and *Brassica oleracea* var. *capitata*. The pharmaceutical composition or the medicinal agent is also used as component of a feedstuff, and the feedstuff containing the pharmaceutical composition or the medicinal agent also exhibit the above-described gastrointestinal function regulating activity.

The pharmaceutical composition or the medicinal agent of the present invention can efficaciously and safely regulate the function(s) of the digestive tract of livestock such as cattle, pigs or horses, poultry such as chicken or Japanese quail, fish such as young yellowtail, sea bream, eel, trout, carp or goldfish, a pet such as a dog or cat, or humans, and is one of the main objects of the present invention.

The term "digestive tract" to be used in this specification refers to organs ranging from the mouth to the anus, particularly to the stomach, duodenum, small intestine, large intestine and rectum.

The term "function(s) of the digestive tract" or "gastrointestinal function(s)" used in this specification means digestion, absorption and digestive motion such as peristalsis. The pharmaceutical composition or the medicinal agent for the digestive tract according to the present invention is useful as a therapeutic and preventive pharmaceutical composition or agent for diarrhea when it acts on the lower part of the digestive tract, such as the large intestine or rectum, while it is efficacious in alleviating a sharp pain and in moderating gastric emptying and borborygmus when it acts on the upper part thereof, such as the stomach or duodenum.

When the pharmaceutical composition or the medicinal agent for regulating the function(s) of the digestive tract is used as a therapeutic and preventive pharmaceutical composition or agent for diarrhea, it is efficacious against various diarrheas which are not limited in cause. Examples of the diarrhea include bacterial diarrhea such as salmonella, viral diarrhea such as caused by adenovirus, parasitic diarrhea such as amebic dysentery, toxic diarrhea such as food and drug poisoning, allergic diarrhea caused by, e.g., food allergy, functional diarrhea such as colic diarrhea and neurotic diarrhea, diarrhea caused by the use of an antibiotic (such as one caused by microbial substitution and staphylococcal diarrhea), and chronic diarrhea caused by the disorder of digestion and absorption, excess gut hormone and colic diseases.

With respect to chickens and pigs, the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is efficacious against bacterial diarrhea such as caused by *Escherichia coli* or swine dysentery, viral diarrhea caused by TGE or rotavirus, or simple diarrhea such as one caused by stress or dietetic diarrhea. Further, it is efficacious also against various diarrheas of fish.

In this case, the dose of *Rosa roxburghii* to be administered varies depending upon the dosage form and the subject animal, so that it is not particularly limited.

For example, an extract of *Rosa roxburghii* is administered to livestock such as a pig in a dose of 20 mg or above, preferably 40 mg or above, still preferably 100 mg or above per kilogram of the body weight.

It will be described that the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for improving antibiotic absorption.

In this case, the pharmaceutical composition or the medicinal agent comprising *Rosa roxburghii* or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. is administered simultaneously with an antibiotic or before or after the administration of an antibiotic. As pharmaceutical composition or the medicinal agent, *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. may be administered by itself in a raw, dried or pulverized state, or a Formulation may be administered. Alternatively, the pharmaceutical composition or the medicinal agent may be added to food or feed in a conventional manner. Namely, the feedstuff comprising a basal diet and least one of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. of the present invention may be used.

The feedstuff of the present invention is prepared by adding *Rosa roxburghii* or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. to the basal diet described above. The amount of *Rosa roxburghii*, or the combination, is preferably 0.001% by weight or above still preferably 0.01% by weight or above in terms of the dried extract thereof, i.e., as the weight of the dried extract thereof.

The pharmaceutical composition, the medicinal agent or the feedstuff for improving antibiotic absorption according to the present invention may be administered simultaneously with an antibiotic or, before or after the administration of an antibiotic. In any case, the effect of improving absorption of an antibiotic can be attained.

In this case, the dose of *Rosa roxburghii* or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. is not particularly limited but varies depending upon the dosage form, the subject animal or the dose of the antibiotic.

For example, when the pharmaceutical composition or the medicinal agent for improving antibiotic absorption comprising two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. is administered to livestock such as a pig a powdered extract, the powdered extract is administered in an amount of part by weight or above, preferably 1 part by weight or above, still preferably 3 parts by weight or above based on 1 part by weight of the antibiotic administered. When *Rosa roxburghii* is administered alone, the amount thereof may be the same as that described above.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for accelerating growth of an animal or improving an egg production rate, weight, egg quality or eggshell strength.

One of the objects of the present invention is to provide a novel pharmaceutical composition, medicinal agent or feedstuff for accelerating growth of an animal, a novel pharmaceutical composition, medicinal agent

or feedstuff for improving the egg production rate, egg weight, egg quality or eggshell strength of animals, and a method for administering them to animals.

More specifically, one of the objects of the present invention is to provide a novel pharmaceutical composition, medicinal agent or feedstuff which can accelerate an increase in the body weight and can improve the survival rate feed conversion ratio of animals, and a novel pharmaceutical composition, medicinal agent or feedstuff which can improve the survival rate, egg production rate, egg weight, egg quality, eggshell strength feed conversion ratio of animals.

The term "growth acceleration" or "accelerating growth of an animal" as used herein includes also growth improvement and growth acceleration of fetuses. The expression "improvement in the egg production rate, egg weight, egg quality and eggshell strength" means an improvement in the production rate, an increase in the egg weight, and improvements in the eggshell and egg quality.

In this case, *Rosa roxburghii* alone or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L., preferably a combination of the three as the active ingredient is administered to animals. *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. can be administered directly to animals or they can be incorporated in the feedstuff or Feed by an ordinary method. Namely, the feedstuff comprises a basal diet and at least one of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. of the present invention may be used.

The feedstuff of the present invention is prepared by adding *Rosa roxburghii* alone or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to the basal diet described above. The amount of *Rosa roxburghii* or the combination is preferably 0.001% by weight or above, still preferably 0.01% by weight or above in terms of the dried extract thereof, i.e., as the weight of the dried extract thereof.

The pharmaceutical composition, the medicinal agent or the feedstuff accelerates the growth of animals, and improves the rate of raising or Feed conversion rate. When it is administered to birds and fish, it accelerates an improvement in the egg production rate, egg weight, egg quality or eggshell strength of them.

The dose of *Rosa roxburghii* is not particularly limited in this case, since it varies depending on the dosage form thereof and the subject animal.

When *Rosa roxburghii* extract is administered to livestock, such as cat and pigs, the amount thereof is usually at least 10 mg, preferably at least 50 mg and still preferably at least 100 mg, based on 1 kilogram of the body weight thereof.

#### DETAILED DESCRIPTION:

#### EXAMPLES

The present invention will now be described in more detail with reference to the following Examples which should not be considered to limit the scope of the present invention.

In the Examples, the description of the dose of each component, e.g., mg/kg p.o." means oral administration in a dose of 10 mg per kilogram body weight. Further, the symbols "\*" and "\*\*" used in the column of t x.sup.2 test in the Tables mean  $p < 0.05$  and  $p < 0.01$ , respectively.

The *Rosa roxburghii* used in Examples A-1, B-1 and B-2 is one commercially available under the trade name of "Rosa roxburghii extract powder MF", which is a product of Maruzen Seiyaku K. K. comprising 30% of an extract of *Rosa roxburghii* and 70% dextrin. The dose of *Rosa roxburghii* is expressed in terms of the weight of this extract powder. The dose of *Artemisiae argyi* folium is expressed in terms of the weight of an extract thereof having 4-fold concentration with respect to its normal weight, while the dose of *Brassica oleracea* var. capitata L. is expressed in terms of the weight of an extract thereof having a 9-fold concentration with respect to its normal weight.

#### EXAMPLE A-1

As shown in Tables A-1 and A-2, *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. were each orally administered (Table A-1), or as a mixture of two or more of them (Table A-2), to the SLC:ICR male mice (age: 5 to 6 weeks, weight: 25 to 33 g) in doses of 2000 mg/kg, while physiological saline was orally administered thereas a control. After 24 hours, clinically available *Escherichia coli* (5.0.times.10.sup.7 CFU/mouse, 0.2 ml) was intravenously inoculated in each mouse to determine the survival rate based on the number of viable mice after 7 days from the infection. The results are given in Tables A-1 and A-2.

TABLE A-1	Survival Sample rate
x.sup.2 test	control (physiological saline p.o.)
0 <i>Artemisiae argyi</i> folium 500 mg/Kg p.o.	20 <i>Artemisiae argyi</i> folium 1000 mg/Kg p.o.
30 <i>Artemisiae argyi</i> folium 2000 mg/Kg p.o.	50 * <i>Brassica oleracea</i> var. capitata L. 500 mg/Kg 10 p.o.
100 <i>Brassica oleracea</i> var. capitata L. 1000 mg/Kg 30 p.o.	20 <i>Brassica oleracea</i> var. capitata L. 2000 mg/Kg 40 *
1000 mg/Kg 40 * p.p.	30 <i>Rosa roxburghii</i> 500 mg/Kg p.o.
2000 mg/Kg 40 *	20 <i>Rosa roxburghii</i> 1000 mg/Kg p.o.
30 <i>Rosa roxburghii</i> 2000 mg/Kg p.o.	50

TABLE A-2	Survival Sample rate
x.sup.2 test	control (physiological saline p.o.)
0 <i>Artemisiae argyi</i> folium 500 mg/Kg p.o.	and 40 * <i>Brassica oleracea</i> var. capitata L. 500 mg/Kg p.o.
500 mg/Kg p.o.	and 70 * <i>Artemisiae argyi</i> folium 1000 mg/Kg p.o.
1000 mg/Kg p.o.	and 100 ** <i>Brassica oleracea</i> var. capitata L. 2000 mg/Kg p.o.
2000 mg/Kg p.o.	<i>Rosa roxburghii</i> 500 mg/Kg p.o.
500 mg/Kg p.o.	and 60 * <i>Artemisiae argyi</i> folium 1000 mg/Kg p.o.
1000 mg/Kg p.o.	<i>Rosa roxburghii</i> 2000 mg/Kg p.o.
2000 mg/Kg p.o.	and 100 ** <i>Artemisiae argyi</i> folium 500 mg/Kg p.o.
500 mg/Kg p.o.	and 70 * <i>Brassica oleracea</i> var. capitata L. 500 mg/Kg p.o.
1000 mg/Kg p.o.	and 100 ** <i>Artemisiae argyi</i> folium 1000 mg/Kg p.o.
2000 mg/Kg p.o.	and <i>Brassica oleracea</i> var. capitata L. 500 mg/Kg p.o.

capitata L. 1000 mg/Kg p.o. \_\_\_\_\_

As shown in Table A-1, the survival rate increases depending upon the of *Rosa roxburghii*, *Artemisiae argyi* folium or *Brassica oleracea* var. capitata L., which reveals that these plants have a protective effect.

As shown in Table A-2, the simultaneous use of *Rosa roxburghii* with *Artemisiae argyi* folium, that of *Artemisiae argyi* folium with *Brassica oleracea* var. capitata L. and that of *Rosa roxburghii* with *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. exhibit the protect effect exceeding the arithmetic sum of the respective effects of these plants. That is, the above simultaneous use exhibits a significant synergistic effect, being of great value.

The above test in Example A-1 reveals that the medicinal agent for immunopotentiating and protecting from infectious diseases of the present invention has an excellent protective activity against Gram-negative bacteria, which suggests that the agent of the present invention is not one which exhibits a low antimicrobial activity only against Gram-positive bacteria like *Artemisiae argyi* folium, but one which potentiates the immune function itself. Accordingly, the agent of the present invention is effective in the prevention and treatment of various maladies and use as a preventive and therapeutic agent for various infectious diseases as a functional food or feed. Thus, the present invention is of great value.

#### EXAMPLE B-1

As shown in Table B-1, *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. were each orally administered to six SLC:SD male rats (age: 6 weeks, weight: 250 to 300 g) in a dose of 125 mg/kg, while physiological saline was orally administered thereto as control. After 30 minutes, 1 ml of castor oil was further orally administered to the rats to determine the time which elapsed until the occurrence of diarrhea and the total amount of excrement given within 24 hours of the administration. The antidiarrheal index of each case was calculated from these values according to the following formula 1 to evaluate the antidiarrheal activity. The measured values and the antidiarrheal index are given in Table B-1. ##EQU1##

TABLE B-1	occurrence amount of Anti-		of diarrhea excrement		Time until the Total	
(g) index	amount	of	diarrhea	excrement	diarrheal	Sample (min)
					control	(physiological)
38.5	+-	5.9	6.7	+-	0.67	1.00 saline p.o.)
13.8	4.5	+-	0.83	2.06	folium 125 mg/Kg p.o.	<i>Artemisiae argyi</i> 53.3
18.4	3.7	+-	1.09	3.38	folium 250 mg/Kg p.o.	<i>Artemisiae argyi</i> 71.8
12.8	2.0	+-	0.68	8.33	folium 500 mg/Kg p.o.	<i>Artemisiae argyi</i> 95.8
+-	7.9	5.9	+-	1.16	1.36 capitata L. 125 mg/Kg p.o.	<i>Brassica oleracea</i> var. 4
49.5	+-	10.7	4.7	+-	1.14	1.83 capitata L. 250 mg/Kg p.o.
51.2	+-	8.2	3.8	+-	1.21	2.34 capitata L. 500 mg/Kg p.o.
50.5	+-	10.0	4.2	+-	1.67	2.09 125 mg/Kg p.o.
68.3	+-	6.4	3.7	+-	0.80	3.21 250 mg/Kg p.o.
77.8	+-	14.4	1.8	+-	0.70	7.52 500 mg/Kg p.o.

It was ascertained from the results given in Table B-1 that *Artemisiae argyi* folium, *Rosa roxburghii* and *Brassica oleracea* var. capitata L. have

antidiarrheal activity.

#### EXAMPLE B-2

As shown in Tables B-2 and B-3, two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. were orally administered to six SLC:SD male rats (age: 6 weeks, weight: 250 to 300 g) each in doses of 125 to 500 mg/kg, while physiological saline was orally administered thereto as a control. After 30 minutes, 1 ml of castor oil was further administered orally to the rats to determine the time elapsed until the occurrence of diarrhea and the total amount of the excrement given within 2 hours of the administration. The antidiarrheal index of each case was calculated from these values according to the above formula 1 to evaluate the antidiarrheal activity. The measured values and the antidiarrheal index are given in Tables B-2 and B-3.

TABLE B-2					Time until the Total
occurrence	amount of Anti-	of diarrhea	excrement	diarrheal	Sample (min
(g) index				control	(physiological
38.5 .+-. 5.9 6.7 .+-. 0.67 1.00	saline p.o.)				
13.4 4.2 .+-. 1.02 2.60	folium 125 mg/Kg p.o. and				
4.54	capitata L. 125 mg/Kg p.o. and				
99.3 .+-. 13.2 1.8 .+-. 0.70 9.59	folium 250 mg/Kg p.o. and				
78.4 .+-. 8.6 3.5 .+-. 1.73 3.89	capitata L. 500 mg/Kg p.o. and				
90.3 .+-. 12.4 2.6 .+-. 1.6.04	500 mg/Kg p.o. and				
101.8 .+-. 14.3 1.6 .+-. 1.58 11.06	500 mg/Kg p.o. and				
	Artemisiae argyi folium 500 mg/Kg p.o.				

TABLE B-3					Time until the Total
occurrence	amount of Anti-	of diarrhea	excrement	diarrheal	Sample (min
(g) index				control	(physiological
38.5 .+-. 5.9 6.7 .+-. 0.67 1.00	saline p.o.)				
.+-. 17.4 3.5 .+-. 1.73 6.37	mg/Kg p.o.,				
102.0 .+-. 13.0 1.8 .+-. 0.70 9.85	mg/Kg p.o.,				
110.8 .+-. 14.0 1.3 .+-. 1.16 14.82	mg p.o.,				
	Artemisiae argyi folium 500 mg/Kg p.o. and				
	Brassica oleracea var capitata L. 500 mg/Kg p.o.				

It has been ascertained from the results given in Tables B-1 and B-2 that the simultaneous use of *Rosa roxburghii* with *Artemisiae argyi* folium exhibits a synergistic antidiarrheal activity, while it has been ascertained from the results given in Tables B-1 and B-3 that the simultaneous use of *Rosa roxburghii* with *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. also exhibits a synergistic antidiarrheal activity.

The above test in Examples B-1 and B-2 reveals that the medicinal agent for regulating a function of a digestive tract according to the present invention has an excellent antidiarrheal activity. Accordingly, the medicinal agent for regulating a function of a digestive tract of the



present invention is effective in the prevention and treatment of diarrhea caused by various diseases, and is useful as a preventive or therapeutic agent for diarrhea or a functional food or feed. Thus, the present invention is of great value.

The mixture of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. used in the following Examples C-1, C-2, D-1 D-6 were prepared as follows:

Raw leaves (3 kg) of *Artemisiae argyi folium* were immersed in 15 l of water and the obtained mixture was boiled for 30 minutes to conduct extraction. The resulting mixture was filtered to recover a filtrate. filtrate was concentrated to 1.8 kg by heating to give an extract. This extract was powdered by spray drying to give a dry powder of the extract of *Artemisiae argyi folium*. Similarly, *Brassica oleracea* var. *capitata* was also converted into a dry powder of the extract thereof. 1500 g of *Rosa roxburghii* extract powder MF (a product of Maruzen Seiyaku K. K. comprising 30% of an extract of *Rosa roxburghii* and 70% of dextrin) was mixed with 675 g of the above dry powder of the extract of *Artemisiae argyi folium* and 225 g of the above dry powder of the extract of *Brassica oleracea* var. *capitata* L. to give 2400 g of a mixed powder (a weight ratio of *Rosa roxburghii*: *Artemisiae argyi folium*: *Brassica oleracea* var. *capitata* L. being 2:3:1).

#### EXAMPLE C-1

The above mixture of the dry powders or each of the dry powders was orally administered to four SLC:ICR male mice (age: 5 to 6 weeks, weight: 25 to 33 g) in a dose given in Table C-1 together with 10 mg per kilogram of body weight of amoxicillin (hereinafter referred to simply as "AMPC") an antibiotic. After 2 hours, blood was gathered from the mice to determine the serum AMPC concentrations, which were compared with those of mice to which AMPC alone had been administered. Each serum AMPC concentration was determined by using a strain of *Sarcina lutea* (Gram-positive and AMPC-sensitive bacterium) according to the disk method (this method of measurement was also employed in Example C-2).

The results are given in Table C-1.

TABLE C-1	Serum AMPC concn.
.mu.g/ml (figures in parentheses being ratio thereof to the concn. of control)	AMPC 10 mg/kg (control)
2.12 +/- 1.14 (1.00)	AMPC 10 mg/kg + a mixed powder of <i>Rosa roxburghii</i>
Artemisiae argyi folium 2.70 +/- 1.22 (1.27)	and <i>Brassica oleracea</i> var. capitata L. 10 mg/kg
AMPC 10 mg/kg + a mixed	3.04 +/- 1.22 (1.43)
powder of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 30 mg/kg	AMPC 10 mg/kg + a mixed
3.11 +/- 1.97 (1.47)	powder of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 90 mg/kg
AMPC 10 mg/kg + a mixed	3.53 +/- 1.93 (1.67)*
powder of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 190 mg/kg	AMPC 10 mg/kg + a mixed
5.62 +/- 0.57 (2.65)**	powder of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 2000 mg/kg
AMPC 10 mg/kg + <i>Rosa</i>	4.87 +/- 1.81 (2.30)
<i>roxburghii</i> 10 mg/kg	AMPC 10 mg/kg + <i>Artemisiae</i>
2.95 +/- 1.13 (1.39)	a folium 10 mg/kg
AMPC 10 mg/kg + <i>Brassica</i>	2.20 +/- 1.11 (1.04)
<i>oleracea</i> var. capitata L. 10 mg/kg	

EXAMPLE C-2

The results are given in Table C-2.

The formulation of the above basal diet are given in Table C-3.

after 0.5 hr	1 hr	2 hrs	3 hrs	6 hrs	
AMPC 10	3.07	6.03	3.71	2.10	0.85 mg/kg (contr
1.55	0.80	0.63	0.40	0.75	AMPC 10 6.03
1.74	0.80	0.63	0.40	0.75	7.81 5.85 3.71 +
(1.58)**	(1.77)*	(2.05)**	of Rosa roxburghii,	Artemisiae argyi folium	
Brassica oleracea var. capitata L.	30 mg/kg				

note) Serum AMPC concentration,  $\mu\text{g/ml}$  The figures in parentheses are the ratios of the serum AMPC concentration to that of the control.

Administration feeding to 5 5 to 7 weeks period weeks of age of age	
Component	Amt. (%) Amt. (%) par
wheat 33.8 20.0 glucose 9.0 5.0 sugar 3.0 -- wheat flour -- 20.0 yellow	
corn -- 23.6 skim milk powder 40.0 10.0 fish meal 3.0 5.0 soybean oil c	
3.0 9.0 Torula yeast 2.0 2.0 soybean oil 4.0 3.0 common salt 0.2 0.2	
tricalcium phosphate 1.2 1.4 mineral mixture 0.2 0.2 vitamin mixture 0	
0.2 flavor, antibiotic and 0.4 0.4 other feed additives	

The results of the above tests suggest that the medicinal agent for improving antibiotic absorption and the feedstuff according to the pre invention are so efficacious in improving the absorption of an antibiotic that the dose of the antibiotic can be reduced. Accordingly, the pre invention is of great value.

40 sound pigs (castrated pigs of about 10 days old; 3.0 to 3.5 kg) wer

divided into four groups, i.e., three groups each consisting of ten pigs to which the mixed powder comprising *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and a control group consisting of ten pigs to which an ordinary basal diet of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered. A feedstuff enriched with the mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. in amounts of 0.03, 0.1 or 0.3% by weight was administered to the pigs of one of the three groups for 30 days. After 30 days, the rate of raising and the feed conversion ratio (feedstuff intake/gain of body weight) were determined to calculate the rate of raising and the feed conversion ratio (feedstuff intake/gain of body weight). The results were compared with those of the control group.

The composition of the artificial milk-based basal diet for piglings was as shown in Table D-1.

The results of the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and the control group were compared with each other. The test was conducted for the gain of body weight in the control group and the groups of the pre-invention. Table D-2 shows the results.

TABLE D-1 Initiation of feeding

5 to 7 weeks	Feeding period	weeks of age	Component	Amt. (%)	A (%)
5.0	5.0	5.0	Sugar	3.0	--
9.0	5.0	5.0	Wheat flour	--	20.0
10.0	10.0	10.0	Yellow corn	--	23.6
40.0	10.0	10.0	Skim milk powder	40.0	10.0
2.0	2.0	2.0	Fish meal	3.0	5.0
1.4	1.4	1.4	Soybean oil cake	3.0	9.0
0.4	0.4	0.4	Torula yeast	2.0	2.0
			Soybean oil	4.0	3.0
			Common salt	0.2	0.2
			Tricalcium phosphate	1	1.4
			Mineral mixture	0.2	0.2
			Vitamin mixture	0.2	0.2
			Flavor, antibiotic		
			and other feed additives		

TABLE D-2 Gain of Feedstuff body

Feed intake (kg/pig)	Weight conversion (A/B)	raising	ratio	Rate of Group (kg/pig)	Test group	Test group	Test group
11.24	5.3	1.33**	2.12	88.8%	(0.03%)	Test group	11.79
1.87	93.0%	(0.1%)	Test group	12.32	5.7	1.04*	2.16
8.50	3.4	1.15	2.50	81.9%	group		

The gain of body weight per pig in the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered was larger than that in the control group, and the feed conversion ratio and the rate of raising were increased in the former group.

#### EXAMPLE D-2

18 bull Holstein calves of about 7 days old were divided into three groups, i.e., two groups each consisting of six calves to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and a control group consisting of six calves to which neither of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered. The mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was mixed into a milk substitute for raising suckling calves and into a feedstuff containing a milk substitute (artificial milk) for raising suckling calves.

in an amount of 0.125 or 0.25% by weight based on the basal diet. The substitute or the feedstuff was repeatedly administered for 20 days. The dose of the mixed powder was 2.5 g/day or 5 g/day per calf. Each calf weighed at the beginning of the test and after the completion of the test (after 20 days) to calculate the gain of body weight and the rate of body weight gain per day.

The composition of the feedstuff containing the milk substitute for raising suckling calves was as shown in Table D-3.

The results of the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and those of the control group are compared with each other. The t-test was conducted in the control group and the groups of the present invention

Table D-4 shows the results.

TABLE D-3				Component Amt. (%)	
				Skim milk powder	60.0
				Dry whey	
Animal fat or oil	20.0	Fish soluble	4.0	Vitamin mixture	0.5
Mineral mixture	0.7				

TABLE D-4				Gain of body weight		Gain of body weight	
Group (kg/calf)		Group (kg/calf/day)					
		Test group		41.0	+-.	7.45**	2.0
+-.	0.29**	2.5 g/day	Test group	42.5	+-.	5.69**	2.13
				+-.	0.27**	5.0	
g/day	Control	25.7	+-.	8.32	1.29	+-.	0.36
				group			

The gain of body weight per day in the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered was larger than that in the control group, and the rate of body weight gain was increased in the former group.

#### EXAMPLE D-3

80 pullet broilers (chunky) were divided into two groups, i.e., a group consisting of 40 broilers to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and a control group consisting of 40 broilers to which neither of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered.

The mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered in an amount of 0.1% by weight based on the basal diet for broilers to the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered for 6 weeks. Each broiler was weighed at the beginning of the test and after 6 weeks. The average gain of body weight in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and that in the control group were calculated.

The composition of the basal diet for broilers was as shown in Table D

The results of the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and those of

control group are compared with each other. The x.sup.2 -test for the average gain of body weight was conducted in the control group and the group to which *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. were administered.

Table D-6 shows the results.

TABLE D-5			Initiation of feeding			
to 5 5 to 6 to 3 weeks weeks of weeks of			Feeding period of age age age			
Component	Amt. (%)	Amt. (%)	Amt. (%)			
Corn	50.45	55.30	55.34	Milo	10.	
15.0 15.01 Soybean oil cake	18.0	13.0	13.01	Rapeseed oil cake	--	2.5 2
Fish meal	8.0	5.0	5.01	Fish soluble	2.0 -- --	Absorbent feed 3.0 -- --
Gluten meal	Alfalfa meal	2.0 -- --	Torula yeast	1.0 -- --	Meat/bone me	
-- 3.0 3.0	Raw rice bran	-- 2.0 2.0	Animal fat or oil	3.3 2.8 2.81	Com	
salt 0.25 0.25 0.25	Calcium carbonate	0.6 0.3 0.30	Dicalcium phosphate			
0.5 0.5 Lysine -- -- --	Methionine	0.18 -- --	Vitamin mixture	0.1 0.1		
Choline chloride	0.05 0.06 0.06	Mineral mixture	0.1 0.1 0.1	Flavor,		
antibiotic and	0.17 0.09 --	other feed additives				

TABLE D-6			Av. body wt. at			
initiation Av. body wt. Av. gain of of feeding after 6 weeks body weig						
Group (g/broiler)	(g/broiler)	(g/broiler)				
			Test group 43.8 .+-. 0.08 2065			
8.7 2021.7 .+-. 8.6** (0.1%)	Control group	43.6 .+-. 0.07 1879 .+-. 10				
1835.4 .+-. 10.1						

The increase in the average body weight in the group to which *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* were administered was larger than that in the control group.

#### EXAMPLE D-4

12,000 young yellowtails having an average body weight of about 600 g divided into two groups, i.e., a group consisting of 6,000 young yellowtails to which *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. were administered and a control group consisting of 6,000 young yellowtails to which neither of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. was administered. mixed powder of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. was mixed with a basal diet for young yellowtails (moist pellets) in an amount of 0.5% by weight based on the basal diet and the resultant mixture was administered to them at intervals of three days for one month. The results were compared with those of the control group. The young yellowtails were raised in a crawl having a surface of about 3 m.times.3 m.times.3 m. During the test period, the feedstuff was administered once a day in an amount of about 3% by weight based on the body weight a day.

The composition of the basal diet for young yellowtails was as shown in Table D-7.

The death rate during the test period in the group to which *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* were administered and that in the control group were compared with each other to evaluate the effect of improving the rate of raising. The x.s

-test was conducted in the control group and the group of the present invention.

The results are given in Table D-8.

TABLE D-7		Component Amt. (%)	
		Fish meal	56.0
		Meat/bone meal	3
Soybean oil cake	5.0	Corn gluten meal	3.0
Torula yeast	2.0	Wheat flour	28.4
Vitamin mixture	1.0	Choline chloride	0.3
Inorg. mixture	1.0	Sodium polyacrylate	0.3

TABLE D-8		No. of No. of died	
samples	samples	Group (fish)	(fish)
Test group (0.1%)	6000	180	** Control group 6000 750

The death rate during the test period in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* were administered was far smaller than that in the control group to prove the effect of improving the rate of raising.

#### EXAMPLE D-5

0.01, 0.03 or 0.10% (based on the amount of the mixed extract powder comprising *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L.), used in a pellet for mice and rats, of a preparation prepared by spraying an aqueous suspension of the mixed extract powder followed by drying, was used to form the pellet for mice and rats. Each feedstuff containing *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. thus prepared was administered to each group male SD rats, which consisted of 8 rats of 21 days old (body weight 50 to 57 g), for one month and then each rat was weighed to calculate rate of body weight gain.

The composition of the basal diet for rats (pellet) was as shown in Table D-9.

The body weights of the rats weighed during the test period in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered were compared with those in the control group to examine the effect on the gain of body weight in both groups. t-test was conducted in the control group and the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata*

The results are given in Table D-10

TABLE D-9		Component Amt. (%)	
		Casein	14.12
		Corn starch	58.40
Glucose	12.83	Soybean oil	5.45
Cellulose	5.00	Minerals	4.00
Vitamins A	0.10	Vitamin B	0.10

#### TABLE D-10

Average body weight (g) and rate of body weight gain in each group Group  
week 1 week 2 weeks 3 weeks 4 weeks

Test group 52.9 .+- . 1.48 106.7 .+- . 4.29 162.4 .+- . 7.17 213.4 .+- . 6  
 258.2 .+- . 7.70 (0.03%) (1.00) (2.02) (3.06) (4.03) (4.86) \* Test grou  
 53.2 .+- . 1.83 107.6 .+- . 3.77 162.6 .+- . 7.28 216.3 .+- . 9.41 258.5 .  
 10.96 (0.1%) (1.00) (2.02) (3.06) (4.07) (4.88) \* Test group 53.7 .+- .  
 2.65 107.6 .+- . 3.23 166.7 .+- . 6.09 221.4 .+- . 10.61 270.2 .+- . 12.53  
 (0.3%) (1.00) (2.00) (3.10) (4.12) (5.03) \*\* Control group 53.8 .+- . 1  
 102.9 .+- . 5.23 157.3 .+- . 9.91 204.5 .+- . 13.41 241.9 .+- . 17.08 (1.0  
 (1.91) (2.92) (3.80) (4.50)

After the administration of *Rosa roxburghii*, *Artemisiae argyi* folium a  
*Brassica oleracea* var. capitata L., an increment in the rate of body  
 weight gain of 0.23 to 0.53 was recognized depending on the dose, as  
 compared with the control group.

#### EXAMPLE D-6

The tests were conducted twice by using groups of Dekalb-TX chickens o  
 182 days old (average body weight: 1.5 kg). Each group consisted of 8  
 chickens. The average results are given in Table D-11 and D-12. In the  
 test, the chickens were divided into three groups, i.e., a group to wh  
 0.08% of the mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium  
*Brassica oleracea* var. capitata L. was administered, a group to which  
 of the mixed powder was administered, and the control group. The feedi  
 was conducted for a period of 18 days.

After the completion of the feeding, body weight and egg weight  
 determination, egg quality test and eggshell strength determination (w  
 an eggshell strength meter) were conducted. Further, the feed conversi  
 ratio (feedstuff intake/gain of body weight) was calculated.

The results are given in Tables D-11 and D-12.

TABLE D-11				Egg Feed prodn. Egg			
weight/day	conversion	Group rate	(g/chicken	multidot.	day)	rate	
				Test group	79.7%	47.3	2.14 (0.0
Test group	79.3%	46.6	2.28 (0.1%)	Control	74.2%	44.2	2.34 group
				Note) Egg weight/day indicates			
total weight of eggs laid by a chicken				a day			

TABLE D-12				Eggshell Eggshell			
Eggshell strength	thickness	amount	Group (kg/cm.sup.2)	(mm)	(wt. %)		
				Test group	3.40	0.358	10.6 (0.0
Test group	3.16	0.359	10.4 (0.1%)	Control	3.12	0.374	10.5 group

It will be apparent from Table D-11 that the egg production rate, egg  
 weight and feed conversion rate were increased more remarkably in the  
 group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica*  
*oleracea* var. capitata L. had been administered than those in the cont  
 group.

It will be apparent from Table D-12 that as compared with the thicknes  
 the eggshell, the eggshell strength was more increased in the group to  
 which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* v  
 capitata L. had been administered than that in the control group. The

results promise the prevention of egg cracking.

One embodiment of the present invention provides a medicinal agent or animal feedstuff for accelerating growth of an animal which comprises roxburghii alone or Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. to be administered to animals directly or as additives to be incorporated into a basal diet so as to improve the body weight gain, survival rate and feed conversion rate of an animal; and another embodiment of the present invention provides a medicinal agent improving an egg production rate, egg weight, egg quality or eggshell strength of animals. Since Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. are used, a high degree of safety is afforded and no environmental pollution is caused. Further, Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata are available at low costs and are excellent as a medicinal agent or animal feedstuff for accelerating growth of an animal and a medicinal agent for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal and have a high storageability.

The invention being thus described, it will be obvious that the same may be varied in many ways. Such variations are not to be regarded as a departure from the spirit and scope of the invention, and all such modifications as would be obvious to one skilled in the art are intended to be included within the scope of the following claims.

#### CLAIMS:

What we claim is:

1. A method of reducing the symptoms of diarrhea in an animal which comprises administering a pharmacologically effective amount of a mixture of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. to the animal.
2. The method of claim 1, wherein the animal is a mammal.
3. The method of claim 2, wherein the mammal is a human being.
4. The method of reducing the symptoms of diarrhea according to claim 1 wherein the mixture is an extract thereof.



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L20: Entry 3 of 3

File: USPT

Sep 17, 1996

US-PAT-NO: 5556624

DOCUMENT-IDENTIFIER: US 5556624 A

TITLE: Method of immunopotentiating and protecting an animal from E. c infections using a combination of Rosa roxburghii, Artemisiae argyi fo and Brassica oleracea var. capitata L.

DATE-ISSUED: September 17, 1996

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APPL-NO: 8/ 333079

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## U.S. PATENT DOCUMENTS

☐ Search Selected☐ Search ALL

PAT-NO	ISSUE-DATE	PATENTEE-NAME	US-CL
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ART-UNIT: 188

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## ABSTRACT:

A method for immunopotentiating and protecting an animal from an infection caused by E. coli is described. The method comprises administering a composition containing Rosa roxburghii, Artemisiae argyi folium, and Brassica oleracea var. capitata L. to the animal.

5. Claims, 0 Drawing figures  
Exemplary Claim Number: 1

## BRIEF SUMMARY:

## BACKGROUND OF THE INVENTION

## 1. Field of the Invention

The present invention relates to a medicinal agent or food which potentiates the immune system of an animal, including a human being, thereby protect the animal from infection, and a feed or feedstuff having an immunopotentiating activity.

The present invention also relates to a medicinal agent or food for regulating a function of the digestive tract or gastrointestinal of an animal, including a human being, and a feed or feedstuff having the activity of regulating the function of the digestive tract or gastrointestinal of an animal.

Further, the present invention relates to a medicinal agent for improving antibiotic absorption by an animal, including a human being, and a feedstuff for improving antibiotic absorption by an animal.

Furthermore, the present invention relates to a medicinal agent for accelerating an animal growth or for improving an egg production rate, weight, egg quality or eggshell strength of an animal and a feed or feedstuff having the activity of accelerating an animal growth or the activity of improving an egg production rate, egg weight, egg quality eggshell strength of an animal.

## 2. Description of the Related Art

With the recent progress in immunology, it has come to be thought that various maladies and infectious diseases of an animal, including a human being, are caused by a weakening or deficiency in the immune system of animal.

For example, a human being frequently suffers from a weakening or deficiency in his immune system because of bronchial asthma, allergic diseases, articular rheumatism, autoimmune diseases, nutritional disorders, surgical operation, aging, cancer, organ transplantation, pregnancy or the like, resulting in the complication of an infectious disease such as respiratory infections, sepsis or urinary infections.

Up to this time, various antibiotics have been administered to patient with such maladies or infectious diseases. Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandry and aquaculture in order to raise livestock, poultry or fish efficiently, and in such raising, it has been also a practice to administer a high dose of an antibiotic.

The repeated administration of an antibiotic for a long period causes generation of antibiotic-resistant bacteria to lower the effect of the antibiotic. Further, hospital infection has also become a significant problem. Under these circumstances, it has been expected to develop a preventive and therapeutic agent which can potentiate the immune system while reducing the dosage of an antibiotic.

Further, the overcrowded raising employed in the fields of animal husbandry and aquaculture has a problem in that various infectious diseases frequently break out because of stress or juvenile immuno-deficiency. Furthermore, when a high dose of an antibiotic is administered as a countermeasure against the problem, there occur other problems that the antibiotic is not completely consumed and that antibiotic-resistant bacteria propagate in the environment.

Recently, animals, including human beings, have frequently suffered from various gastrointestinal diseases which are caused by a lowering or deficiency in the immune system, stress, dyspepsia or the like, and most of which are accompanied with diarrhea.

For example, a human being becomes susceptible to an infectious disease of the digestive tract as his resistance lowers, and representative examples of the infectious disease include bacterial, viral and parasitic diarrheas. Further, the above gastro-intestinal diseases also include acute diarrheas caused by food poisoning and food allergy and chronic diarrheas caused by a disorder of digestion and absorption, excess gut hormone and colic diseases.

It has been the practice to administer an intestinal depressomotor for intestines, an astringent, an irritant-absorbing agent, a torpent for enteric mucous membranes or various antibiotics against these gastrointestinal diseases.

Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandry and aquaculture to raise livestock, poultry or fish efficiently. In such raising, it has been a practice to administer the above therapeutic agents for the treatment of diarrhea caused by stress or dyspepsia. Particularly, a large amount of an antibiotic has been used for the prevention of infectious diseases.

The administration of the above depressomotor or astringent is essentially a nosotropic means, while that of an antibiotic is an etiotropic one. However, the repeated administration of an antibiotic for the prevention of infectious diseases has a problem that the preventive effect lowers owing to the generation of antibiotic-resistant bacteria.

Under these circumstances, it has been expected to develop a novel preventive and therapeutic agent which can regulate gastrointestinal functions themselves and is safe.

Up to this time, various antibiotics have been administered to patient with various maladies or infectious diseases. Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandry and aquaculture to raise livestock, poultry or fish efficiently. In raising, it has been a practice to administer a high dose of an antibiotic.

With respect to the use of an antibiotic, however, there have been problems such that the antibiotic accumulates in the body of an animal to affect the body and that the administration of a more potent antibiotic becomes necessary because of the generation of antibiotic-resistant bacteria. Further, the administration of an antibiotic to an animal is problematic also in that the environment is polluted with the antibiotic contained in the excrement of the animal. Under these circumstances, it has been expected that the dose of the antibiotic to be administered is reduced. In order to reduce the dose of the antibiotic, it has been believed effective that the absorption of the antibiotic through the digestive tract should be enhanced to attain an effect equal or superior to that attained by the administration thereof in a high dose, even when the antibiotic is administered in a low dose. However, no substance which can improve the absorption of an antibiotic has been found as yet.

In preparing animal feed, various attempts have been widely made at improving the feeding efficiency for animals and also at accelerating growth thereof by adding various antibiotics to the feed, incorporating increased amount of proteins into the feed, changing the feeding method, improving the dosage form of the feed.

With respect to the use of an antibiotic, however, there have been problems such that the antibiotic remains or accumulates in the body of an animal to affect the body and that the administration of a more potent antibiotic becomes necessary because of the generation of resistant bacteria. Further, the administration of an antibiotic to an animal is problematic also in that the environment is polluted with the antibiotic contained in the excrement of the animal. In addition, the effects

obtained by varying the feeding method and the dosage form of the feed limited. Under these circumstances, it is necessary to develop a safe less problematic animal growth accelerator, feed and improver for the production ratio, egg weight, egg quality or eggshell strength of animal.

## DISCLOSURE OF THE INVENTION

### Summary of the Invention

In view of the above problems, the present inventors have extensively studied for many years and have found that *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. are useful for overcoming the problems. The present invention has been accomplished on the basis of this finding.

Thus, the present invention provides;

(1) a pharmaceutical composition comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(2) a pharmaceutical composition comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.;

(3) a feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.;

(4) a feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a basal diet;

(5) a method for feeding a feedstuff comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to an animal; and

(6) a method for feeding a feedstuff comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a basal diet to an animal.

In view of the above problems with respect to the use of antibiotics, present inventors have extensively studied for many years on protective agents which are safe for animals and have found that *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. have an immunopotentiating activity. The present invention has been accomplished on the basis of this finding.

Thus, the present invention provides;

(7) a method for immunopotentiating and protecting from infectious diseases, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.

to an animal;

(8) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for preparing a medicine for an animal for immunopotentiating and protecting from infectious diseases;

(9) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for immunopotentiating and protecting from infectious diseases in an animal;

(10) a pharmaceutical composition for immunopotentiating and protecting from infectious diseases in an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* and a pharmaceutically acceptable carrier;

(11) a medicinal agent for immunopotentiating and protecting from infectious diseases of an animal, comprising a substance selected from group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier; a

(12) a feedstuff for an animal, which is useful for immunopotentiating protecting from infectious diseases in an animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.

In view of the above problem, the present inventors have extensively studied for many years regulating agents for the digestive tract which safe for human beings and animals, and have found that *Rosa roxburghii* *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. have the activity of regulating gastrointestinal functions. The present invention has been accomplished on the basis of this finding.

Thus, the present invention provides;

(13) a method for regulating the digestive tract, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* *Brassica oleracea* var. *capitata* L. to an animal;

(14) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for preparing a medicine for an animal for regulating the digestive tract

(15) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for regulating the digestive tract of an animal;

(16) a pharmaceutical composition for regulating the digestive tract of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(17) a medicinal agent for regulating the digestive tract of an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* and a pharmaceutically acceptable carrier; and

(18) a feedstuff for an animal, which is useful for regulating the digestive tract of the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.

To provide an absorbefacient which enables a reduction in the dose of antibiotic when administered together with the antibiotic, the present inventors have extensively studied to find out that the absorption of antibiotic can surprisingly be improved when *Rosa roxburghii* alone or or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. are administered simultaneously with the antibiotic or before or after the administration of the antibiotic.

Thus, the present invention provides;

(19) a method for improving antibiotic absorption, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* *Brassica oleracea* var. *capitata* L. to an animal;

(20) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for preparing a medicine for an animal for improving its antibiotic absorption;

(21) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for improving antibiotic absorption in an animal;

(22) a pharmaceutical composition for improving antibiotic absorption in an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(23) a medicinal agent for improving antibiotic absorption in an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* and a pharmaceutically acceptable carrier; and

(24) a feedstuff for an animal, which is useful for improving antibiotic absorption in the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.

To provide a feedstuff free from the abovementioned defects of an ordinary feedstuff and a method of administering the feedstuff to animals, the present inventors have extensively studied. As a result, they have found that the growth of animals, including fetuses, is surprisingly accelerated by administering *Rosa roxburghii* alone or two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea*

var. capitata L. to them. The inventors have further found that an improvement in the egg production rate, egg weight, egg quality or eggshell strength of birds and fishes is also accelerated by them.

Thus, the present invention provides;

(25) a method for accelerating growth of an animal, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium *Brassica oleracea* var. capitata L. to the animal;

(26) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata for preparing a medicine for an animal for accelerating growth of the animal;

(27) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata for accelerating growth of an animal;

(28) a pharmaceutical composition for accelerating the growth of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. and a pharmaceutically acceptable carrier;

(29) a medicinal agent for accelerating the growth of an animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata and a pharmaceutically acceptable carrier;

(30) a feedstuff for an animal, which is useful for accelerating the growth of the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L.;

(31) a method for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. to the animal;

(32) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata for preparing a medicine of an animal for improving an egg production rate, egg weight, egg quality or eggshell strength of the animal;

(33) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal;

(34) a pharmaceutical composition for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. and a pharmaceutically acceptable carrier;



(35) a medicinal agent for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier; and

(36) a feedstuff for an animal, which is useful for improving an egg production rate, egg weight, egg quality or eggshell strength of the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata*

Further scope and applicability of the present invention will become apparent from the detailed description given hereinafter. However, it should be understood that the detailed description and specific examples while indicating preferred embodiments of the invention, are given by way of illustration only, since various changes and modifications within the spirit and scope of the invention will become apparent to those skilled in the art from this detailed description.

#### DETAILED DESCRIPTION OF THE INVENTION

*Rosa roxburghii* is a perennial shrub of the family Rosaceae and is native to Guizhou in China and its fruit has been noticed as the material of juice, jam or liquor. The fruit of *Rosa roxburghii* has been known to have a pharmacological activity and is useful as an antiulcer agent by virtue of its cancer-preventing, cholesterol level-lowering and antistress activities.

*Artemisiae argyi* folium is a plant of the family Compositae and has been known to be useful as an antidiarrheal or antiabdominalgia agent, a hemostatic or the like. Further, this plant has been known to exhibit low antimicrobial activity only against Gram-positive bacteria. *Artemisiae argyi* folium includes, for example, *Artemisia princeps* Pampanini, *Artemisia mongolica* Fischer, *Artemisia argyi* LEVL. et VANT., and *Artemisia lavandulaefolia* DC.

*Brassica oleracea* var. *capitata* L. is a plant of the family Cruciferae and has been used as food.

The *Rosa roxburghii* to be used in the present invention is not particularly limited in form, but may have any form so far as it contains the essences of *Rosa roxburghii*. Generally, the fruit of *Rosa roxburghii* may be used in its raw state or as a dry powder prepared by conventional means prior to use, or an extract prepared by using water, an organic solvent or a mixture of both as an extractant.

The *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. to be used in the present invention are not particularly limited in form, but may have any form so far as it contains the essences thereof. Generally, the leaf of *Artemisiae argyi* folium or *Brassica oleracea* var. *capitata* may be used in a raw state or as a dry powder prepared by conventional means prior to use, or an extract prepared by using water, an organic solvent or a mixture of both as an extractant.

An extract from the above leaves can be prepared by, e.g., a process with

comprises immersing 1 part by weight of the raw material in 5 parts by weight of water, boiling the obtained mixture under heating for 30 min to conduct extraction, filtering the resulting system, and concentrating the obtained filtrate to 3.6 parts by weight. The obtained extract may be powdered by spray drying, freeze drying, vacuum drying (vacuum concentration) or the like.

When an organic solvent is used for extraction, methanol, ethanol, n-propanol, n-butanol, acetone, ethyl acetate, ether, methylene chloride, chloroform, benzene, carbon tetrachloride and petroleum ether are preferable. These organic solvents may be used alone or as a mixture of two or more of them.

The extracts thus produced may be used as such or may be concentrated, diluted or freed from the solvent prior to use.

The extract of *Rosa roxburghii* to be used in the present invention may be one commercially available under the trade name of "Rosa roxburghii extract powder MF", which is a product of Maruzen Seiyaku K.K. comprising 30% of an extract of *Rosa roxburghii* and 70% of dextrin.

In the present invention, at least one of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. is(are) used.

When *Rosa roxburghii* and *Artemisiae argyi* folium are used in the present invention, the ratio between them is not particularly limited. Generally, when *Artemisiae argyi* folium is used, in terms of raw leaf, that is, when it is prescribed as the weight of raw leaf, in an amount of 0.25 to 400 parts by weight, preferably 0.5 to 200 parts by weight, still preferably 1 to 10 parts by weight, based on 1 part by weight of the extract of *Rosa roxburghii*.

When both *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* are used together with *Rosa roxburghii*, the amounts thereof are not particularly limited. Generally, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. are used in amounts of 0.25 to 400 parts by weight and 0.5 to 800 parts by weight, respectively, in terms of their respective raw leaf, based on 1 part by weight of the extract of *Rosa roxburghii*. It is preferable that *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. be used in amounts of 0.5 to 200 parts by weight and 1 to 400 parts by weight respectively, still preferably in amounts of 1 to 100 parts by weight and 5 to 200 parts by weight respectively.

The pharmaceutical composition or the medicinal agent of the present invention may be administered, with the purpose of the prevention of diseases, as a food having a regulating effect on a living body, i.e., so-called functional food, which can be prepared by adding this composition or agent to food.

When the pharmaceutical composition or the medicinal agent of the present invention comprising one or more members selected from among *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* is used as a medicine or so-called health food, it may be in the form of a tablet, granule, powder, capsule or syrup. The pharmaceutical composition or the medicinal agent is prepared by mixing the members with a conventional filler or carrier, binder, lubricant or the like and treating the obtained mixture in the conventional manner.

When the pharmaceutical composition or the medicinal agent of the present invention comprising one or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* is used for an animal, such as a mammal, a bird and a fish, especially a livestock, the dosage form thereof is not particularly limited. For example, it may be administered to livestock in a state mixed with a basal diet (or feed or feedstuff). That is, this composition or agent may be mixed with a basal diet (or feed or feedstuff) just before using or may be premixed with a basal diet (or feed or feedstuff). In other words, the feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. of the present invention may be administered to an animal as a feed having a biophylactic and regulating function.

The arbitrary basal diet for animals, which is used for preparing the feedstuff according to the present invention, is not particularly limited. Examples of the raw materials constituting the basal diet include grains such as corn, milo and wheat flour, brans such as defatted rice bran and wheat bran, animal substances such as fish meal and skim milk, vegetable oil cake such as soybean oil cake, and additives such as calcium carbonate, calcium phosphate, common salt, vitamin B<sub>2</sub>, DL-methionine, choline chloride, manganese sulfate, dry iron sulfate, calcium iodate, copper sulfate, dry zinc sulfate and sodium saccharin. The basal diet may be prepared by blending some members selected from among these materials. The formulation of the basal diet varies depending upon the animal to which the diet is administered.

The improved feed of the present invention, i.e., the feedstuff containing at least one of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. can be administered to various livestock, poultry, pets and fish. The livestock includes pig, cattle, horse, sheep and rabbit; the poultry includes chicken, Japanese quail, turkey and duck; the pets include dog and cat; and the fish include yellowtail, sea bream, flatfish, globefish, hardtail, amberjack, salmon, carp, eel, sweetfish, trout, and lobster.

Needless to say, the pharmaceutical composition, the medicinal agent and the feedstuff of the present invention are non-toxic.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for immunopotentiating and protecting from infectious diseases.

In this case, *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. may be used alone or as a combination of two or more of them. In other words, the combination of *Rosa roxburghii* with *Artemisiae argyi folium*, that of *Rosa roxburghii* with *Brassica oleracea* var. *capitata* L., that of *Artemisiae argyi folium* with *Brassica oleracea* var. *capitata* L., and that of *Rosa roxburghii* with *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. can be used.

*Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. exhibit an immunopotentiating and protective activity even when used each alone. However, the use of a combination of two or more of them surprisingly exhibits an immunopotentiating and protective activity which is higher than that of the use of each of them.

The pharmaceutical composition or the medicinal agent of the present invention can safely and effectively potentiate the immune function of livestock such as cattle, pig or horse, a poultry such as chicken or Japanese quail, fish such as young yellowtail, sea bream, eel, trout, or goldfish, a pet such as dog or cat, or a human being to protect the animal or the like from various infectious diseases, which is one of the objects of the present invention.

The term "immunopotential" used in this specification means "potentiation of the immune function of an animal such as a human being, mammal, fish or the like".

The pharmaceutical composition or the medicinal agent of the present invention have the effect of enhancing the immune function of an animal including a human being, and serves as a preventive and therapeutic pharmaceutical composition or agent for various maladies and infectious diseases by virtue of this effect, so that the diseases against which the composition or agent is efficacious are not particularly limited. For example, it is efficacious against, e.g., articular rheumatism, autoimmune diseases, bronchial asthma, nutritional disorders, surgical operation, diseases of old age and various infectious diseases such as respiratory infection, sepsis and urinary infection with respect to a human being.

With respect to animals other than humans, this composition or agent is efficacious against scours, epizootic pneumonia, atrophic rhinitis and infectious enterogastroenteritis of a pig, pneumonia and Marek's disease of chicken, scours, pneumonia and mastitis of cattle, and AIDS and leukemia of a pet.

Further, with respect to fish, the infectious disease against which the composition or agent of the present invention is efficacious is not particularly limited and includes bacterial diseases such as streptococcosis and nodosity, and viral diseases.

In this case, the dose of *Rosa roxburghii* to be administered varies depending upon the dosage form and the subject animal, so that it is not particularly limited.

For example, an extract of *Rosa roxburghii* is administered to livestock such as a pig in a dose of 25 mg or above, preferably 50 mg or above, still preferably 100 mg or above per kilogram of the body weight.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for regulating the digestive tract.

In this case, *Rosa roxburghii* is preferably used as the essential component. The activity of regulating the digestive tract, i.e., the gastrointestinal activity, of the pharmaceutical composition or the medicinal agent is synergistically enhanced when the composition or agent further contains *Artemisiae argyi folium* or *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. The pharmaceutical composition or the medicinal agent is also used as a component of a feedstuff, and the feedstuff containing the pharmaceutical composition or the medicinal agent also exhibits the above-described gastrointestinal function regulating activity.

The pharmaceutical composition or the medicinal agent of the present invention can efficaciously and safely regulate the function(s) of the digestive tract of livestock such as cattle, pig or horse, poultry such as chicken or Japanese quail, fish such as young yellowtail, sea bream, eel, trout, carp or goldfish, a pet such as a dog or cat, or humans, which are one of the main objects of the present invention.

The term "digestive tract" to be used in this specification refers to organs ranging from the mouth to the anus, particularly to the stomach, duodenum, small intestine, large intestine and rectum.

The term "function(s) of the digestive tract" or "gastrointestinal function(s)" used in this specification means digestion, absorption and digestive motion such as peristalsis. The pharmaceutical composition or the medicinal agent for the digestive tract according to the present invention is useful as a therapeutic and preventive pharmaceutical composition or agent for diarrhea when it acts on the lower part of the digestive tract, such as the large intestine or rectum, while it is efficacious in alleviating a sharp pain and in moderating gastric emptying and borborygmus when it acts on the upper part thereof, such as the stomach or duodenum.

When the pharmaceutical composition or the medicinal agent for regulating the function(s) of the digestive tract is used as a therapeutic and preventive pharmaceutical composition or agent for diarrhea, it is efficacious against various diarrheas which are not limited in cause. Examples of the diarrhea include bacterial diarrhea such as salmonella, viral diarrhea such as caused by adenovirus, parasitic diarrhea such as amebic dysentery, toxic diarrhea such as food and drug poisoning, allergic diarrhea caused by, e.g., food allergy, functional diarrheas such as celiac diarrhea and neurotic diarrhea, diarrheas caused by the use of an antibiotic (such as one caused by microbial substitution and staphylococcal diarrhea), and chronic diarrheas caused by the disorder of digestion and absorption, excess gut hormone and colic diseases.

With respect to chickens and pigs, the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is efficacious against bacterial diarrhea such as caused by *Escherichia coli* or swine dysentery, viral diarrhea caused by TGE or rotavirus, or simple diarrhea such as one caused by stress or dietetic diarrhea. Further, it is efficacious also against various diarrheas of fish.

In this case, the dose of *Rosa roxburghii* to be administered varies depending upon the dosage form and the subject animal, so that it is not particularly limited.

For example, an extract of *Rosa roxburghii* is administered to livestock such as a pig in a dose of 20 mg or above, preferably 40 mg or above, still preferably 100 mg or above per kilogram of the body weight.

It will be described about the case that the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for improving antibiotic absorption.

In this case, the pharmaceutical composition or the medicinal agent comprising *Rosa roxburghii* or a combination of two or more members

selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. is administered simultaneously with an antibiotic or before or after the administration of an antibiotic. As pharmaceutical composition or the medicinal agent, *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. may be administered by itself in a raw, dried or pulverized state, or a formulation may be administered. Alternatively, the pharmaceutical composition or the medicinal agent may be added to food or feed in a conventional manner. Namely, the feedstuff comprising a basal diet and least one of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. of the present invention may be used.

The feedstuff of the present invention is prepared by adding *Rosa roxburghii* or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to the basal diet described above. The amount of *Rosa roxburghii*, or the combination, is preferably 0.001% by weight or above still preferably 0.01% by weight or above in terms of the dried extract thereof, i.e., as the weight of the dried extract thereof.

The pharmaceutical composition, the medicinal agent or the feedstuff for improving antibiotic absorption according to the present invention may be administered simultaneously with an antibiotic or, before or after the administration of an antibiotic. In any case, the effect of improving absorption of an antibiotic can be attained.

In this case, the dose of *Rosa roxburghii* or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. is not particularly limited but varies depending upon the dosage form, the subject animal or the dose of the antibiotic.

For example, when the pharmaceutical composition or the medicinal agent for improving antibiotic absorption comprising two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. is administered to livestock such as a pig a powdered extract, the powdered extract is administered in an amount of part by weight or above, preferably 1 part by weight or above, still preferably 3 parts by weight or above based on 1 part by weight of the antibiotic administered. When *Rosa roxburghii* is administered alone, the amount thereof may be the same as that described above.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for accelerating growth of an animal or improving an egg production rate, weight, egg quality or eggshell strength.

One of the objects of the present invention is to provide a novel pharmaceutical composition, medicinal agent or feedstuff for accelerating growth of an animal, a novel pharmaceutical composition, medicinal agent or feedstuff for improving the egg production rate, egg weight, egg quality or eggshell strength of animals, and a method for administering them to animals.

More specifically, one of the objects of the present invention is to provide a novel pharmaceutical composition, medicinal agent or feedstuff which can accelerate an increase in the body weight and can improve the

survival rate, feed conversion ratio of animals, and a novel pharmaceutical composition, medicinal agent or feedstuff which can improve the survival rate, egg production rate, egg weight, egg quality, eggshell strength and feed conversion ratio of animals.

The term "growth acceleration" or "accelerating growth of an animal" as used herein includes also growth improvement and growth acceleration of fetuses. The expression "improvement in the egg production ratio, egg weight, egg quality and eggshell strength" means an improvement in the production ratio, an increase in the egg weight, and improvements in the eggshell and egg quality.

In this case, *Rosa roxburghii* alone or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L., preferably a combination of three members described above, as the active ingredient is administered to animals. *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. can be administered directly to animals or they can be incorporated in the feedstuff or feed by an ordinary method. Namely, the feedstuff comprising a basal diet and at least one of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. of the present invention may be used.

The feedstuff of the present invention is prepared by adding *Rosa roxburghii* alone or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to the basal diet described above. The amount of *Rosa roxburghii* or the combination is preferably 0.001% by weight or above, still preferably 0.01% by weight or above in terms of the dried extract thereof, i.e., as the weight of the dried extract thereof.

The pharmaceutical composition, the medicinal agent or the feedstuff accelerates the growth of animals, and improves the rate of raising or feed conversion rate. When it is administered to birds and fishes, it accelerates an improvement in the egg production rate, egg weight, egg quality or eggshell strength of them.

The dose of *Rosa roxburghii* is not particularly limited in this case, since it varies depending on the dosage form thereof and the subject animal.

When *Rosa roxburghii* extract is administered to livestock, such as cat and pigs, the amount thereof is usually at least 10 mg, preferably at least 50 mg and still preferably at least 100 mg, based on 1 kilogram of the body weight thereof.

#### DETAILED DESCRIPTION:

#### EXAMPLES

The present invention will now be described in more detail with reference to the following Examples which should not be considered to limit the scope of the present invention.

In the Examples, the description of the dose of each component, e.g., mg/kg p.o." means oral administration in a dose of 10 mg per kilogram

body weight. Further, the symbols "\*" and "\*\*\*" used in the column of t x.sup.2 test in Tables mean  $p < 0.05$  and  $p < 0.01$ , respectively.

The *Rosa roxburghii* used in Examples A-1, B-1 and B-2 is one commercial available under the trade name of "Rosa roxburghii extract powder MF", which is a product of Maruzen Seiyaku K K comprising 30% of an extract *Rosa roxburghii* and 70% dextrin. The dose of *Rosa roxburghii* is expressed in terms of the weight of this extract powder. The dose of *Artemisiae argyi* folium is expressed in terms of the weight of an extract thereof having 4-fold concentration with respect to its normal weight, while the dose of *Brassica oleracea* var. *capitata* L. is expressed in terms of the weight of an extract having a 9-fold concentration with respect to its normal weight.

#### Example A-1

As shown in Tables A-1 and A-2, *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were each orally administered a (Table A-1), or as a mixture of two or more of them (Table A-2), to the SLC:ICR male mice (age: 5 to 6 weeks, weight: 25 to 33 g) in doses of to 2000 mg/kg, while physiological saline was orally administered there as a control. After 24 hours, clinically available *Escherichia coli* (5.0.times.10.sup.7 CFU/mouse, 0.2 ml) was intravenously inoculated in each mouse to determine the survival rate based on the number of viable mice after 7 days from the infection. The results are given in Tables A-1 and A-2.

TABLE A-1	x.sup.2 Sample Survival rate (%) test	control
(physiological saline p.o.)	0	<i>Artemisiae argyi</i> folium 500 mg/Kg p.o. 2
<i>Artemisiae argyi</i> folium 1000 mg/Kg p.o.	30	<i>Artemisiae argyi</i> folium 200 mg/Kg p.o. 50 *
<i>Brassica oleracea</i> var. <i>capitata</i> L. 10	500 mg/Kg p.o.	<i>Brassica oleracea</i> var. <i>capitata</i> L. 30
1000 mg/Kg p.o.	<i>Brassica oleracea</i> var. <i>capitata</i> L. 40 *	2000 mg/Kg p.p.
<i>Rosa roxburghii</i> 500 mg/Kg p.o.	2	<i>Rosa roxburghii</i> 1000 mg/Kg p.o.
30	<i>Rosa roxburghii</i> 2000 mg/Kg p.o.	50

TABLE A-2	Survival x.sup.2 Sample rate (%) test	control
(physiological saline p.o.)	0	<i>Artemisiae argyi</i> folium 500 mg/Kg p.o. a
40 *	<i>Brassica oleracea</i> var. <i>capitata</i> L. 500 mg/Kg p.o.	<i>Artemisiae argyi</i> folium 1000 mg/Kg p.o. and 70 *
<i>Brassica oleracea</i> var. <i>capitata</i> L. 100	mg/Kg p.o.	<i>Artemisiae argyi</i> folium 2000 mg/Kg p.o. and 100 **
<i>Brassica oleracea</i> var. <i>capitata</i> L. 2000 mg/Kg p.o.	<i>Rosa roxburghii</i> 500 mg/Kg p.	and 60 *
<i>Artemisiae argyi</i> folium 500 mg/Kg p.o.	<i>Rosa roxburghii</i> 1000 mg p.o. and 90 **	<i>Artemisiae argyi</i> folium 1000 mg/Kg p.o.
<i>Rosa roxburghii</i> 2000 mg/Kg p.o. and 100 **	<i>Artemisiae argyi</i> folium 2000 mg/Kg p.o.	<i>Rosa roxburghii</i> 500 mg/Kg p.o., 70 *
<i>Artemisiae argyi</i> folium 500 mg/Kg p.o.	<i>Brassica oleracea</i> var. <i>capitata</i> L. 500 mg/Kg p.o.	<i>Rosa roxburghii</i> 1000 mg/Kg p.o., 100 **
<i>Artemisiae argyi</i> folium 1000 mg/Kg p.o.	and <i>Brassica oleracea</i> var. <i>capitata</i> L. 1000 mg/Kg p.o.	

As shown in Table A-1, the survival rate increases depending upon the of *Rosa roxburghii*, *Artemisiae argyi* folium or *Brassica oleracea* var. *capitata* L., which reveals that these plants have a protective effect.



As shown in Table A-2, the simultaneous use of *Rosa roxburghii* with *Artemisiae argyi* folium, that of *Artemisiae argyi* folium with *Brassica oleracea* var. *capitata* L. and that of *Rosa roxburghii* with *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. exhibit the protect effect exceeding the arithmetic sum of the respective effects of these plants. That is, the above simultaneous use exhibits a significant synergistic effect, being of great value.

The above test in Example A-1 reveals that the medicinal agent for immunopotentiating and protecting from infectious diseases of the present invention has an excellent protective activity against Gram-negative bacteria, which suggests that the agent of the present invention is not one which exhibits a low antimicrobial activity only against Gram-positive bacteria like *Escherichia coli*, but one which potentiates the immune function itself. Accordingly, the agent of the present invention is effective in the prevention and treatment of various maladies and use as a preventive and therapeutic agent for various infectious diseases as a functional food or feed. Thus, the present invention is of great value.

#### Example B-1

As shown in Table B-1, *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were each orally administered to six SLC:SD male rats (age: 6 weeks, weight: 250 to 300 g) in a dose of 125 500 mg/kg, while physiological saline was orally administered thereto as control. After 30 minutes, 1 ml of castor oil was further orally administered to the rats to determine the time which elapsed until the occurrence of diarrhea and the total amount of excrement given within 24 hours of the administration. The antidiarrheal index of each case was calculated from these values according to the following formula 1 to evaluate the antidiarrheal activity. The measured values and the antidiarrheal index are given in Table B-1. ##EQU1##

TABLE B-1	Time until the Total	
occurrence amount of Anti- of diarrhea excrement	diarrheal Sample (min	control (physiological
(g) index		
38.5 .+- . 5.9 6.7 .+- . 0.67 1.00	saline p.o.)	<i>Artemisiae argyi</i> 53.3 .+
13.8 4.5 .+- . 0.83 2.06	folium 125 mg/Kg p.o.	<i>Artemisiae argyi</i> 71.8 .+
18.4 3.7 .+- . 1.09 3.38	folium 250 mg/Kg p.o.	<i>Artemisiae argyi</i> 95.8 .+
12.8 2.0 .+- . 0.68 8.33	folium 500 mg/Kg p.o.	<i>Brassica oleracea</i> var. 4
+- . 7.9 5.9 .+- . 1.16 1.36	<i>capitata</i> L. 125 mg/Kg p.o.	<i>Brassica olerac</i>
var. 49.5 .+- . 10.7 4.7 .+- . 1.14 1.83	<i>capitata</i> L. 250 mg/Kg p.o.	var. 51.2 .+- . 8.2 3.8 .+- . 1.21 2.34
<i>capitata</i> L. 500 mg/Kg p		
<i>Rosa roxburghii</i> 50.5 .+- . 10.0 4.2 .+- . 1.67 2.09	125 mg/Kg p.o.	<i>Rosa</i>
<i>roxburghii</i> 68.3 .+- . 6.4 3.7 .+- . 0.80 3.21	250 mg/Kg p.o.	<i>Rosa roxbur</i>
77.8 .+- . 14.4 1.8 .+- . 0.70 7.52	500 mg/Kg p.o.	

It was ascertained from the results given in

Table B-1 that *Artemisiae argyi* folium, *Rosa roxburghii* and *Brassica oleracea* var. *capitata* L. had an antidiarrheal activity.

#### Example B-2

As shown in Tables B-2 and B-3, two or more members selected from among

Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were orally administered to six SLC:SD male rats (age: 6 weeks, weight: 250 to 300 g) each in doses of 125 to 500 mg/kg, while physiological saline was orally administered thereto as a control. After 30 minutes, 1 ml of castor oil was further administered orally to the rats to determine the time elapsed until the occurrence of diarrhea and the total amount of the excrement given within 2 hours of the administration. The antidiarrheal index of each case was calculated from these values according to the above formula 1 to evaluate the antidiarrheal activity. The measured values and the antidiarrheal index are given in Tables B-2 and B-3.

TABLE B-2									Time until the Total
occurrence	amount of Anti-	of diarrhea	excrement	diarrheal	Sample (min				control (physiologica
(g) index									
38.5	+-	5.9	6.7	+-	0.67	1.00	saline p.o.)	Artemisiae argyi folium	
62.7	+-	13.4	4.2	+-	1.02	2.60	mg/Kg p.o. and	Brassica oleracea var	
capitata L.							125 mg/Kg p.o.	Artemisiae argyi folium	250 83.5
	+-	1.45	4.54	mg/Kg p.o. and			Brassica oleracea var.	capitata L.	250 m
p.o. Artemisiae argyi folium				500	99.3	+-	13.2	1.8	+-
p.o. and Brassica oleracea var.				capitata L.	500	mg/Kg p.o.	Rosa roxbur		
78.4	+-	8.6	3.5	+-	1.73	3.89	125 mg/Kg p.o. and	Artemisiae argyi	
folium 125 mg/Kg p.o.				Rosa roxburghii	90.3	+-	12.4	2.6	+-
250 mg/Kg p.o. and Artemisiae argyi folium				250 mg/Kg p.o.			Rosa roxburg		
500 mg/Kg p.o. and				101.8	+-	14.3	1.6	+-	1.58
folium 500 mg/Kg p.o.									11.06
									Artemisiae argy

TABLE B-3									Time until the Total
occurrence	amount of Anti-	of diarrhea	excrement	diarrheal	Sample (min				control (physiologica
(g) index									
38.5	+-	5.9	6.7	+-	0.67	1.00	saline p.o.)	Rosa roxburghii 125	91.5
	+-	17.4	3.5	+-	1.73	6.37	mg/Kg p.o.,	Artemisiae argyi folium 125 m	
p.o. and Brassica oleracea var.				capitata L.	125	mg/Kg p.o.	Rosa roxbur		
250	102.0	+-	13.0	1.8	+-	0.70	9.85	mg/Kg p.o.,	Artemisiae argyi fo
250 mg/Kg p.o. and Brassica oleracea var.				capitata L.	250	mg/Kg p.o.	R		
roxburghii 500 mg/Kg p.o.,				Artemisiae argyi folium	500	mg/Kg	110.8	+-	
14.0	1.3	+-	1.16	14.82	p.o. and	Brassica oleracea var.	capitata L.	5	
mg/Kg p.o.									

It has been ascertained from the results given in Tables B-1 and B-2 that the simultaneous use of Rosa roxburghii with Artemisiae argyi folium exhibits a synergistic antidiarrheal activity, while it has been ascertained from the results given in Tables B-1 and B-3 that the simultaneous use of Rosa roxburghii with Artemisiae argyi folium and Brassica oleracea var. capitata L. also exhibits a synergistic antidiarrheal activity.

The above test in Examples B-1 and B-2 reveals that the medicinal agent for regulating a function of a digestive tract according to the present invention has an excellent antidiarrheal activity. Accordingly, the medicinal agent for regulating a function of a digestive tract of the present invention is effective in the prevention and treatment of diarrhea caused by various diseases, and is useful as a preventive or therapeutic agent for diarrhea or a functional food or feed. Thus, the present invention is of great value.

The mixture of Rosa roxburghii, Artemisiae argyi folium and Brassica

oleracea var. capitata L. used in the following Examples C-1, C-2, D-1 D-6 were prepared as follows:

Raw leaves (3 kg) of *Artemisiae argyi folium* were immersed in 15 l of water and the obtained mixture was boiled for 30 minutes to conduct extraction. The resulting mixture was filtered to recover a filtrate. filtrate was concentrated to 1.8 kg by heating to give an extract. This extract was powdered by spray drying to give a dry powder of the extract of *Artemisiae argyi folium*. Similarly, *Brassica oleracea* var. capitata was also converted into a dry powder of the extract thereof. 1500 g of *Rosa roxburghii* extract powder MF (a product of Maruzen Seiyaku K.K. comprising 30% of an extract of *Rosa roxburghii* and 70% of dextrin) was mixed with 675 g of the above dry powder of the extract of *Artemisiae argyi folium* and 225 g of the above dry powder of the extract of *Brassica oleracea* var. capitata L. to give 2400 g of a mixed powder (a weight ratio of *Rosa roxburghii*:*Artemisiae argyi folium*:*Brassica oleracea* var. capitata L. being 2:3:1).

#### Example C-1

The above mixture of the dry powders or each of the dry powders was orally administered to four SLC:ICR male mice (age: 5 to 6 weeks, weight: 25 to 33 g) in a dose given in Table C-1 together with 10 mg per kilogram of body weight of amoxicillin (hereinafter referred to simply as "AMPC") an antibiotic. After 2 hours, blood was gathered from the mice to determine the serum AMPC concentrations, which were compared with those of mice to which AMPC alone had been administered. Each serum AMPC concentration was determined by using a strain of *Sarcina lutea* (Gram-positive and AMPC-sensitive bacterium) according to the disk method (this method of measurement was also employed in Example C-2).

The results are given in Table C-1.

TABLE C-1	Serum AMPC concn.
.mu.g/ml (figures in parentheses being ratio thereof to the concn. of control)	AMPC 10 mg/kg (control)
2.12 +/- 1.14 (1.00) AMPC 10 mg/kg + a mixed	2.70 +/- 1.22 (1.27) po
of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 10 mg/kg	AMPC 10 mg/kg + a mixed
3.04 +/- 1.22 (1.43) powder of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 30 mg/kg	AMPC 10 mg/kg + a mixed
3.11 +/- 1.97 (1.47) powder of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 90 mg/kg	AMPC 10 mg/kg + a mixed
3.53 +/- 1.93 (1.67)* powder of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 190 mg/kg	AMPC 10 mg/kg + a mixed
5.62 +/- 0.57 (2.65)** powder of <i>Rosa roxburghii</i> , <i>Artemisiae argyi folium</i> and <i>Brassica oleracea</i> var. capitata L. 2000 mg/kg	AMPC 10 mg/kg + <i>Rosa</i>
4.87 +/- 1.81 (2.30) <i>roxburghii</i> 10 mg/kg	AMPC 10 mg/kg + <i>Artemisiae</i>
2.95 +/- 1.13 (1.39) <i>folium</i> 10 mg/kg	AMPC 10 mg/kg + <i>Brassica</i>
2.20 +/- 1.11 (1.04) <i>oleracea</i> var. capitata L. 10 mg/kg	

As shown in Table C-1, the group of the mice to which AMPC and *Rosa roxburghii* had been administered and the groups of the mice to which *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. capitata L. had been administered were apparently improved in the absorption of AMPC depending upon the dose thereof as compared with the group of the mice to which AMPC alone had been administered.

## Example C-2

AMPC (10 mg/kg) alone or the mixture of AMPC (10 mg/kg) with the mixed powder (30 mg/kg) comprising *Rosa roxburghii*, *Artemisiae argyi* folium *Brassica oleracea* var. *capitata* L. was administered to six SPF male pi (age: 1 month, weight: about 15 kg) mixed with a basal diet to gather blood from the pigs after 0.5, 1, 2, 3 and 6 hours. The serum AMPC concentrations of the pigs to which AMPC and the mixed dry powder had administered were determined and compared with those of the pigs to wh AMPC alone had been administered.

The results are given in Table C-2.

The formulation of the above basal diet are given in Table C-3.

TABLE C-2

	after	after	after	after	after	0.5 hr	1 hr	2 hrs	3 hrs	6 hrs
AMPC 10 mg/kg (control)	3.07	+-	1.55	6.03	+-	0.80	3.71	+-	0.63	2
+- 0.40 0.85 +- 0.75	AMPC 10 mg/kg + a mixed powder of	6.03	+-	1						
7.81 +- 1.49 5.85 +- 0.80	3.71 +- 1.41	1.74 +- 1.16	Rosa							
roxburghii, <i>Artemisiae argyi</i> (1.63)**	(1.30)*	(1.58)**	(1.77)*	(2.05)*						
folium and <i>Brassica oleracea</i> var. <i>capitata</i> L.	30 mg/kg									

note) Serum AMPC concentration, .mu.g/ml The figures in parentheses are the ratios of the serum AMPC concentration to that of the control.

TABLE C-3				Initiation of							
Administration feeding to 5 5 to 7 weeks period weeks of age of age											
Component	Amt. (%)	Amt. (%)		par							
wheat	33.8	20.0	glucose	9.0	5.0	sugar	3.0	--	wheat flour	--	20.0
corn	--	23.6	skimmilk powder	40.0	10.0	fish meal	3.0	5.0	soybean oil	c	
	3.0	9.0	Torula yeast	2.0	2.0	soybean oil	4.0	3.0	common salt	0.2	0.2
			tricalcium phosphate	1.2	1.4	mineral mixture	0.2	0.2	vitamin mixture	0	
			0.2 flavor, antibiotic and	0.4	0.4	other feed additives					

As shown in Table C-2, the pigs to which AMPC and the mixed powder had been administered were apparently improved in the absorption of AMPC o a period of 0.5 to 6 hours after the administration as compared with t pigs to which AMPC alone had been administered.

The results of the above tests suggest that the medicinal agent for improving antibiotic absorption and the feedstuff according to the pre invention are so efficacious in improving the absorption of an antibio that the dose of the antibiotic can be reduced. Accordingly, the prese invention is of great value.

## Example D-1

40 sound pigs (castrated pigs of about 10 days old; 3.0 to 3.5 kg) wer divided into four groups, i.e., three groups each consisting of ten pi to which the mixed powder comprising *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and a control group consisting of ten pigs to which an ordinary basal diet f

of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. was administered. A feedstuff enriched with the mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. in amounts of 0.03, 0.1 or 0.3% by weight was administered to the pigs of one of the three groups for 30 days. After 30 days, the intake of the feedstuff and the gain of body weight were determined to calculate the rate of raising and the feed conversion ratio (feedstuff intake/gain of body weight). The results were compared with those of the control group.

The composition of the artificial milk-based basal diet for piglings was as shown in Table D-1.

The results of the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. were administered and the control group were compared with each other. The t-test was conducted on the gain of body weight in the control group and the groups of the prevention. Table D-2 shows the results.

TABLE D-1 \_\_\_\_\_ Initiation of feeding

5 to 7 weeks	Feeding period	weeks of age	Component	Amt. (%)	A
5.0	5.0	5.0	Sugar	3.0	--
9.0	5.0	5.0	Wheat flour	20.0	Yellow corn
40.0	10.0	10.0	Fish meal	3.0	5.0
2.0	2.0	2.0	Soybean oil	4.0	3.0
1.4	0.2	0.2	Mineral mixture	0.2	0.2
0.4	0.2	0.2	Vitamin mixture	0.2	0.2
			Flavor, antibiotic		
			and other feed additives		

TABLE D-2 \_\_\_\_\_ Feedstuff Gain of Feed

intake (kg/pig)	body weight (kg/pig)	conversion (A/B)	raising	(A) (B)	ratio	Rate of Group (kg/pig)
11.24	5.3	1.33**	2.12	88.8%	(0.03%)	Test group
1.87	93.0%	(0.1%)	Test group	12.32	5.7	1.04*
8.50	3.4	1.15	2.50	81.9%	group	

The gain of body weight per pig in the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. were administered was larger than that in the control group, and the feed conversion ratio and the rate of raising were increased in the former group.

#### Example D-2

18 bull Holstein calves of about 7 days old were divided into three groups, i.e., two groups each consisting of six calves to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. were administered and a control group consisting of six calves to which neither of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. was administered. The mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. was mixed into a milk substitute for raising suckling calves and into a feedstuff containing a milk substitute (artificial milk) for raising suckling calves in an amount of 0.125 or 0.25% by weight based on the basal diet. The substitute or the feedstuff was repeatedly administered for 20 days. The dose of the mixed powder was 2.5 g/day or 5 g/day per calf. Each calf was weighed at the beginning of the test and after the completion of the test.

(after 20 days) to calculate the gain of body weight and the rate of body weight gain per day.

The composition of the feedstuff containing the milk substitute for raising suckling calves was as shown in Table D-3.

The results of the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and those of the control group are compared with each other. The t-test was conducted in the control group and the groups of the present invention

Table D-4 shows the results.

TABLE D-3				Component Amt. (%)	
				Skim milk powder	60.0
				Dry whey	
Animal fat or oil	20.0	Fish soluble	4.0	Vitamin mixture	0.5
mixture	0.7			Mineral	

TABLE D-4				Gain of body weight			
weight		Group (kg/calf)		(kg/calf/day)			
				Test group		41.0	
.+-. 0.29**		2.5 g/day		Test group		42.5	
g/day		Control		25.7		.+-. 8.32	
						1.29	
						.+-. 0.36	
						group	

The gain of body weight per day in the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered was larger than that in the control group, and the rate of body weight gain was increased in the former group.

#### Example D-3

80 pullet broilers (chunky) were divided into two groups, i.e., a group consisting of 40 broilers to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and a control group consisting of 40 broilers to which neither of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered.

The mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered in an amount of 0.1% by weight based on the basal diet for broilers to the group, to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered, for 6 weeks. Each broiler was weighed at the beginning of the test and after 6 weeks. The average gain of body weight in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and that in the control group were calculated.

The composition of the basal diet for broilers was as shown in Table D

The results of the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and those of the control group are compared with each other. The x<sup>2</sup>-test for the average gain of body weight was conducted in the control group and the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered.

Table D-6 shows the results.

TABLE D-5				Initiation of feeding			
to 5 5 to 6 to 3 weeks weeks of weeks of Feeding period of age age age							
Component	Amt. (%)	Amt. (%)	Amt. (%)				
				Corn	50.45	55.30	55.34
15.0	15.01	Soybean oil cake	18.0	13.0	13.01	Rapeseed oil cake	-- 2.5
Fish meal	8.0	5.0	5.01	Fish soluble	2.0	-- --	Absorbent feed
Gluten meal	3.0	3.0	Raw rice bran	-- 2.0	2.0	Animal fat or oil	3.3 2.8 2.81
Alfalfa meal	2.0	-- --	Torula yeast	1.0	-- --	Meat/bone me	
Calcium carbonate	0.6	0.3	0.30	Dicalcium phosphate			
Lysine	-- --	-- --	Methionine	0.18	-- --	Vitamin mixture	0.1 0.1
Choline chloride	0.05	0.06	0.06	Mineral mixture	0.1	0.1	0.1
antibiotic and	0.17	0.09	--	other feed additives			

TABLE D-6				Av. body wt. at			
initiation Av. body wt. Av. gain of of feeding after 6 weeks body weig							
Group (g/broiler)	(g/broiler)	(g/broiler)	(g/broiler)				
				Test group	43.8	+-.	0.08
8.7	2021.7	+-.	8.6** (0.1%)	Control group	43.6	+-.	0.07
1835.4	+-.	10.1					

The increase in the average body weight in the group to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata were administered was larger than that in the control group.

#### Example D-4

12,000 young yellowtails having an average body weight of about 600 g divided into two groups, i.e., a group consisting of 6,000 young yellowtails to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were administered and a control group consisting of 6,000 young yellowtails to which neither of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. was administered. mixed powder of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were mixed with a basal diet for young yellowtails (Moist pellets) in an amount of 0.5% by weight based on the basal diet and the resultant mixture was administered to them at intervals of three days for one month. The results were compared with those of the control group. The young yellowtails were raised in a crawl having a size of about 3 m.times.3 m.times.3 m. During the test period, the feedstuff was administered once a day in an amount of about 3% by weight based on the body weight a day.

The composition of the basal diet for young yellowtails was as shown in Table D-7.

The death rate during the test period in the group to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata were administered and that in the control group were compared with each other to evaluate the effect of improving the rate of raising. The x.s.-test was conducted in the control group and the group of the present invention.

The results are given in Table D-8.

TABLE D-7		Component Amt. (%)			
		Fish meal		56.0	Meat/bone meal 3
Soybean oil cake	5.0	Corn gluten meal	3.0	Torula yeast	2.0
28.4	Vitamin mixture	1.0	Choline chloride	0.3	Inorg. mixture 1.0
polyacrylate	0.3				

TABLE D-8		No. of No. of died	
samples	samples	Group (fish)	(fish)
Test group (0.1%)	6000	180**	control group 6000 750

The death rate during the test period in the group to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata were administered was far smaller than that in the control group to prove the effect of improving the rate of raising.

#### Example D-5

0.01, 0.03 or 0.10%, based on the amount of the mixed extract powder comprising Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. used in a pellet for mice and rats, of a preparation prepared by spraying an aqueous suspension of the mixed extract powder followed by drying, was used to form the pellet for mice and rats. Each feedstuff containing Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. thus prepared was administered to each group male slc:SD rats, which consisted of 8 rats of 21 days old (body weight 50 to 57 g), for one month and then each rat was weighed to calculate rate of body weight gain.

The composition of the basal diet for rats (pellet) was as shown in Table D-9.

The body weights of the rats weighed during the test period in the group to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were administered were compared with those in the control group to examine the effect on the gain of body weight in both groups. A t-test was conducted in the control group and the groups to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata

The results are given in Table D-10

TABLE D-9		Component Amt. (%)			
		Casein		14.12	Corn starch 58.40
Glucose	12.83	Soybean oil	5.45	Cellulose	5.00
D 0.10	Vitamin B 0.10	Minerals	4.00	Vitamins A	

TABLE D-10										Average body weight			
and rate of body weight gain in each group										Group 0	week 1	week 2	weeks
weeks 4 weeks										Test group 52.9			
106.7	+-	162.4	+-	213.4	+-	258.2	+-	(0.03%)	1.48	4.29	7.17	6.9	
7.70	(1.00)	(2.02)	(3.06)	(4.03)	(4.86)*	Test group 53.2				+-	107.6	+	
162.6	+-	216.3	+-	258.5	+-	(0.1%)	1.83	3.77	7.28	9.41	10.96	(1.0	
(2.02)	(3.06)	(4.07)	(4.88)*	Test group 53.7				+-	107.6	+-	166.7	+-	
221.4	+-	270.2	+-	(0.3%)	2.65	3.23	6.09	10.61	12.53	(1.00)	(2.00)		
(3.10)	(4.12)	(5.03)**	Control 53.8				+-	102.9	+-	157.3	+-	204.5	



241.9 .+-. group 1.38 5.23 9.91 13.41 17.08 (1.00) (1.91) (2.92) (3.80  
(4.50)

After the administration of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L., an increment in the rate of body weight gain of 0.23 to 0.53 was recognized depending on the dose, as compared with the control group.

#### Example D-6

The tests were conducted twice by using groups of Dekalb-TX chickens 182 days old (average body weight: 1.5 kg). Each group consisted of 8 chickens. The average results are given in Table D-11 and D-12. In the test, the chickens were divided into three groups, i.e., a group to which 0.03% of the mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered, a group to which 0.1% of the mixed powder was administered, and the control group. The feeding was conducted for a period of 18 days.

After the completion of the feeding, body weight and egg weight determination, egg quality test and eggshell strength determination (with an eggshell strength meter) were conducted. Further, the feed conversion ratio (feedstuff intake/gain of body weight) was calculated.

The results are given in Tables D-11 and D-12.

TABLE D-11	Egg Feed prodn. Egg			
weight/day	conversion	Group rate	(g/chicken .multidot. day)	ratio
Test group	79.3%	46.6	2.28 (0.1%)	Control 74.2% 44.2 2.34 group
Note) Egg weight/day indicates total weight of eggs laid by a chicken a day				

TABLE D-12	Eggshell Eggshell			
strength thickness	Eggshell	Group	(kg/cm.sup.2) (mm)	(wt. %)
Test group	3.16	0.359	10.4 (0.1%)	Control 3.12 0.374 10.5 group

It will be apparent from Table D-11 that the egg production rate, egg weight and feed conversion ratio were increased more remarkably in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. had been administered than those in the control group.

It will be apparent from Table D-12 that as compared with the thickness of the eggshell, the eggshell strength was more increased in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. had been administered than that in the control group. The results promise the prevention of egg cracking.

One embodiment of the present invention provides a medicinal agent or animal feedstuff for accelerating growth of an animal which comprises *roxburghii* alone or *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. to be administered to animals directly or as additives to be incorporated into a basal diet so as to improve the body weight gain, survival rate and feed conversion ratio of an animal; and

another embodiment of the present invention provides a medicinal agent improving an egg production rate, egg weight, egg quality or eggshell strength of animals. Since *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. are used, a high degree of safety is afforded and no environmental pollution is caused. Further, *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* are available at low costs and are excellent as a medicinal agent or animal feedstuff for accelerating growth of an animal and a medicinal agent for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal and have a high storageability.

The invention being thus described, it will be obvious that the same may be varied in many ways. Such variations are not to be regarded as a departure from the spirit and scope of the invention, and all such modifications as would be obvious to one skilled in the art are intended to be included within the scope of the following claims.

#### CLAIMS:

What we claim is:

1. A method for immunopotentiating and protecting an animal from an infection caused by *Escherichia coli*, which comprises administering a pharmaceutically effective amount of a mixture of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* L. to the animal.
2. The method of claim 1, wherein said infection is selected from the group consisting of a respiratory infection, sepsis and a urinary infection.
3. The method for immunopotentiating and protecting from an infection according to claim 1, wherein the mixture is an extract thereof.
4. The method for immunopotentiating and protecting from an infection according to claim 1, wherein the animal is a mammal.
5. The method for immunopotentiating and protecting from an infection according to claim 4, wherein the mammal is a human.

**WEST****End of Result Set**

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L21: Entry 1 of 1

File: USPT

Apr 29, 1997

US-PAT-NO: 5624671DOCUMENT-IDENTIFIER: US 5624671 A

TITLE: Method for increasing egg production rate, egg weight or eggshe strength by administering a composition containing the plants Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitat

DATE-ISSUED: April 29, 1997

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APPL-NO: 8/ 438494

DATE FILED: May 10, 1995

## PARENT-CASE:

This is a division of Ser. No. 08/390 243, filed Feb. 17, 1995, which division of Ser. No. 08/333 079, filed Nov. 1, 1994, now U.S. Pat. No. 5,556,624 which is a division of Ser. No. 08/285 980, filed Aug. 4, 1993, now abandoned, which is a division of Ser. No. 08/133 708, filed Oct. 1993, now abandoned.

## FOREIGN-APPL-PRIORITY-DATA:

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426/807

PRIOR-ART-DISCLOSED:

U.S. PATENT DOCUMENTS

	Search Selected	Search ALL	
PAT-NO	ISSUE-DATE	PATENTEE-NAME	US-CL
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<input type="checkbox"/> 4197294	April 1980	Klein	424/195
<input type="checkbox"/> 5085871	February 1992	Horikawa et al.	426/2
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ART-UNIT: 188

PRIMARY-EXAMINER: Wityshyn; Michael G.

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ATTY-AGENT-FIRM: Flynn, Thiel, Boutell & Tanis, P.C.

ABSTRACT:

A pharmaceutical composition or feed, which is useful for immunopotentiating and protecting an animal from infectious diseases, regulating a function of the digestive tract, improving antibiotic absorption, accelerating growth of an animal or improving egg production rate, egg weight, egg quality or eggshell strength of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi* folium *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier.

6 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

BRIEF SUMMARY:

## BACKGROUND OF THE INVENTION

### 1. Field of the Invention

The present invention relates to a medicinal agent or food which potentiates the immune system of an animal, including a human being, thereby protect the animal from infection, and a feed or feedstuff have an immunopotentiating activity.

The present invention also relates to a medicinal agent or food for regulating a function of the digestive tract or gastrointestinal of an animal, including a human being, and a feed or feedstuff having the activity of regulating the function of the digestive tract or gastrointestinal of an animal.

Further, the present invention relates to a medicinal agent for improving antibiotic absorption by an animal, including a human being, and a feedstuff for improving antibiotic absorption by an animal.

Furthermore, the present invention relates to a medicinal agent for accelerating animal growth or for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal and a feed or feedstuff having the activity of accelerating an animal growth or the activity of improving an egg production rate, egg weight, egg quality eggshell strength of an animal.

### 2. Description of the Related Art

With the recent progress in immunology, it has come to be thought that various maladies and infectious diseases of an animal, including a human being, are caused by a weakening or deficiency in the immune system of animal.

For example, a human being frequently suffers from a weakening or deficiency in his immune system because of bronchial asthma, allergic diseases, articular rheumatism, autoimmune diseases, nutritional disorders, surgical operation, aging, cancer, organ transplantation, pregnancy or the like, resulting in the complication of a disease such as respiratory infections, sepsis or urinary infections.

Up to this time, various antibiotics have been administered to patients with such maladies or infectious diseases. Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandry and aquaculture in order to raise livestock, poultry or fish efficiently and in such raising, it has been also a practice to administer a high dose of an antibiotic.

The repeated administration of an antibiotic for a long period causes generation of antibiotic resistant bacteria to lower the effect of the antibiotic. Further, hospital infection has also become a significant problem. Under these circumstances, it has been expected to develop a preventive and therapeutic agent which can potentiate the immune system while reducing the dosage of an antibiotic.

Further, the overcrowded raising employed in the fields of animal

husbandry and aquaculture has a problem in that various infectious diseases frequently break out because of stress or juvenile immunodeficiency. Furthermore, when a high dose of an antibiotic is administered as a countermeasure against the problem, there occur other problems that the antibiotic is not completely consumed and antibiotic resistant bacteria propagate in the environment.

Recently, animals, including human beings, have frequently suffered from various gastrointestinal diseases which are caused by a lowering or deficiency in the immune system, stress, dyspepsia or the like, and most of which are accompanied with diarrhea.

For example, a human being becomes susceptible to an infectious disease of the digestive tract as his resistance lowers, and representative examples of the infectious diseases include bacterial, viral and parasitic diarrheas. Further, the above gastrointestinal diseases also include diarrheas caused by food poisoning and food allergy and chronic diarrhea caused by a disorder of digestion and absorption, excess gut hormone and colic diseases.

It has been the practice to administer an intestinal depressomotor for intestines, an astringent, an irritant-absorbing agent, a sorbent for enteric mucous membranes or various antibiotics against these gastrointestinal diseases.

Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandry and aquaculture to raise livestock, poultry and fish efficiently. In such raising, it has been a practice to administer the above therapeutic agents for the treatment of diarrhea caused by stress or dyspepsia. Particularly, a large amount of an antibiotic has been used for the prevention of infectious diseases.

The administration of the above depressomotor or astringent is essentially a nosotropic means, while that of an antibiotic is an etiotropic one. However, the repeated administration of an antibiotic for the prevention of infectious diseases has a problem that the preventive effect is low owing to the generation of antibiotic-resistant bacteria.

Under these circumstances, it has been expected to develop a novel preventive and therapeutic agent which can regulate gastrointestinal functions themselves and is safe.

Up to this time, various antibiotics have been administered to patients with various maladies or infectious diseases. Meanwhile, large-scale or overcrowded raising has been employed in the fields of animal husbandry and aquaculture to raise livestock, poultry and fish efficiently. In raising, it has been a practice to administer a high dose of an antibiotic.

With respect to the use of an antibiotic, however, there have been problems such that the antibiotic accumulates in the body of an animal to affect the body and that the administration of a more potent antibiotic becomes necessary because of the generation of antibiotic-resistant bacteria. Further, the administration of an antibiotic to an animal is problematic also in that the environment is polluted with the antibiotic contained in the excrement of the animal. Under these circumstances, it has been expected that the dose of the antibiotic to be administered is

reduced. In order to reduce the dose of the antibiotic, it has been believed effective that the absorption of the antibiotic through the digestive tract should be enhanced to attain an effect equal or superior to that attained by the administration thereof in a high dose, even when the antibiotic is administered in a low dose. However, no substance which can improve the absorption of an antibiotic has been found as yet.

In preparing animal feed, various attempts have been widely made at improving the feeding efficiency for animals and also at accelerating growth thereof by adding various antibiotics to the feed, incorporating increased amount of proteins into the feed, changing the feeding method, improving the dosage form of the feed.

With respect to the use of an antibiotic, however, there have been pointed out problems such that the antibiotic remains or accumulates in the body of an animal to affect the body and that the administration of a more potent antibiotic becomes necessary because of the generation of resistant bacteria. Further, the administration of an antibiotic to an animal is problematic also in that the environment is polluted with the antibiotic contained in the excrement of the animal. In addition, the effects obtained by varying the feeding method and the dosage form of the feed are limited. Under these circumstances, it is necessary to develop a safe and less problematic animal growth accelerator, feed and improver for the production rate, egg weight, egg quality or eggshell strength of animals.

#### DISCLOSURE OF THE INVENTION

##### SUMMARY OF THE INVENTION

In view of the above problems, the present inventors have extensively studied for many years and have found that *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. are useful for overcoming the problems. The present invention has been accomplished on the basis of this finding.

Thus, the present invention provides:

(1) a pharmaceutical composition comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(2) a pharmaceutical composition comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.;

(3) a feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L.;

(4) a feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a basal diet;

(5) a method for feeding a feedstuff comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and

Brassica oleracea var. capitata L. to an animal; and

(6) a method for feeding a feedstuff comprising a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. and a basal diet to an animal.

In view of the above problems with respect to the use of antibiotics, present inventors have extensively studied for many years on protective agents which are safe for animals and have found that Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. have an immunopotentiating activity. The present invention has been accomplished on the basis of this finding.

Thus, the present invention provides;

(7) a method for immunopotentiating and protecting from infectious diseases, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. to an animal;

(8) a use of a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. for preparing a medicine for an animal for immunopotentiating and protecting from infectious diseases;

(9) a use of a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. for immunopotentiating and protecting from infectious diseases in an animal;

(10) a pharmaceutical composition for immunopotentiating and protecting from infectious diseases in an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. and a pharmaceutically acceptable carrier;

(11) a medicinal agent for immunopotentiating and protecting from infectious diseases of an animal, comprising a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. and a pharmaceutically acceptable carrier; and

(12) a feedstuff for an animal, which is useful for immunopotentiating and protecting from infectious diseases in an animal, comprising a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L..

In view of the above problem, the present inventors have extensively studied for many years regulating agents for the digestive tract which are safe for human beings and animals, and have found that Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. have the activity of regulating gastrointestinal functions. The present invention has been accomplished on the basis of this finding.

Thus, the present invention provides;

(13) a method for regulating the digestive tract, which comprises



administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to an animal;

(14) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for preparing a medicine for an animal for regulating the digestive tract;

(15) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for regulating the digestive tract of an animal;

(16) a pharmaceutical composition for regulating the digestive tract of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(17) a medicinal agent for regulating the digestive tract of an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* and a pharmaceutically acceptable carrier; and

(18) a feedstuff for an animal, which is useful for regulating the digestive tract of the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L..

To provide an absorbent which enables a reduction in the dose of antibiotic when administered together with the antibiotic, the present inventors have extensively studied to find out that the absorption of antibiotic can surprisingly be improved when *Rosa roxburghii* alone or or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. are administered simultaneously with the antibiotic or before or after the administration of the antibiotic.

Thus, the present invention provides;

(19) a method for improving antibiotic absorption, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to an animal;

(20) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for preparing a medicine for an animal for improving its antibiotic absorption;

(21) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for improving antibiotic absorption in an animal;

(22) a pharmaceutical composition for improving antibiotic absorption in an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi*

folium and Brassica oleracea var. capitata L. and a pharmaceutically acceptable carrier;

(23) a medicinal agent for improving antibiotic absorption in an animal comprising a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata and a pharmaceutically acceptable carrier; and

(24) a feedstuff for an animal, which is useful for improving antibiotic absorption in the animal, comprising a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L..

To provide a feedstuff free from the above-mentioned defects of an ordinary feedstuff and a method of administering the feedstuff to an animal, the present inventors have extensively studied. As a result, they have found that the growth of animals, including fetuses, is surprisingly accelerated by administering Rosa roxburghii alone or two or more members selected from among Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. to them. The inventors have further found that an improvement in the egg production rate, egg weight, egg quality or eggshell strength of birds and fish is also accelerated by them.

Thus, the present invention provides;

(25) a method for accelerating growth of an animal, which comprises administering a pharmacologically effective amount of a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. to the animal;

(26) a use of a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata for preparing a medicine for an animal for accelerating growth of the animal;

(27) a use of a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata for accelerating growth of an animal;

(28) a pharmaceutical composition for accelerating the growth of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. and a pharmaceutically acceptable carrier;

(29) a medicinal agent for accelerating the growth of an animal, comprising a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata and a pharmaceutically acceptable carrier;

(30) a feedstuff for an animal, which is useful for accelerating the growth of the animal, comprising a substance selected from the group consisting of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L.;

(31) a method for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal, which comprises administering

pharmacologically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. to the animal;

(32) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for preparing a medicine of an animal for improving an egg production rate, egg weight, egg quality or eggshell strength of the animal;

(33) a use of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal;

(34) a pharmaceutical composition for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal, comprising a pharmaceutically effective amount of a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier;

(35) a medicinal agent for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. and a pharmaceutically acceptable carrier; and

(36) a feedstuff for an animal, which is useful for improving an egg production rate, egg weight, egg quality or eggshell strength of the animal, comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L..

Further scope and applicability of the present invention will become apparent from the detailed description given hereinafter. However, it should be understood that the detailed description and specific examples while indicating preferred embodiments of the invention, are given by way of illustration only, since various changes and modifications within the spirit and scope of the invention will become apparent to those skilled in the art from this detailed description.

#### DETAILED DESCRIPTION OF THE INVENTION

*Rosa roxburghii* is a perennial shrub of the family Rosaceae and is native to Guizhou in China and its fruit has been used as the material of jui jam or liquor. The fruit of *Rosa roxburghii* has been known to have a pharmacological activity and is useful as an antiulcer agent by virtue of its cancer-preventing, cholesterol level-lowering and antistress activities.

*Artemisiae argyi folium* is a plant of the family Compositae and has been known to be useful as an antidiarrheal or antiabdominalgia agent, a hemostatic or the like. Further, this plant has been known to exhibit low antimicrobial activity only against Gram-positive bacteria. *Artemisiae argyi folium* includes, for example, *Artemisia princeps* Pampanini, *Artemisia mongolica* Fischer, *Artemisia argyi* LEVL. et VANT., and *Artemisia lavandulaefolia* DC.

*Brassica oleracea* var. *capitata* L. is a plant of the family *Cruciferae* has been used as food.

The *Rosa roxburghii* to be used in the present invention is not particularly limited in form, but may have any form so far as it contains the essences of *Rosa roxburghii*. Generally, the fruit of *Rosa roxburghii* may be used in its raw state or as a dry powder prepared by conventional means prior to use, or an extract prepared by using water, an organic solvent or a mixture of both as an extractant.

The *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. to be used in the present invention are not particularly limited in form, but may have any form so far as it contains the essences thereof. Generally, the leaf of *Artemisiae argyi* folium or *Brassica oleracea* var. *capitata* may be used in a raw state or as a dry powder prepared by a conventional means prior to use, or an extract prepared by using water, an organic solvent or a mixture of both as an extractant.

An extract from the above leaves can be prepared by, e.g., a process which comprises immersing 1 part by weight of the raw material in 5 parts by weight of water, boiling the obtained mixture under heating for 30 min to conduct extraction, filtering the resulting system, and concentrating the obtained filtrate to 3.6 parts by weight. The obtained extract may be powdered by spray drying, freeze drying, vacuum drying (vacuum concentration) or the like.

When an organic solvent is used for extraction, methanol, ethanol, n-propanol, n-butanol, acetone, ethyl acetate, ether, methylene chloride, chloroform, benzene, carbon tetrachloride and petroleum ether are preferable. These organic solvents may be used alone or as a mixture of two or more of them.

The extracts thus produced may be used as such or may be concentrated, diluted or freed from the solvent prior to use.

The extract of *Rosa roxburghii* to be used in the present invention may be one commercially available under the trade name of "Rosa roxburghii extract powder MF", which is a product of Maruzen Seiyaku K. K. comprising 30% of an extract of *Rosa roxburghii* and 70% of dextrin.

In the present invention, at least one of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. is(are) used.

When *Rosa roxburghii* and *Artemisiae argyi* folium are used in the present invention, the ratio between them is not particularly limited. Generally, when *Artemisiae argyi* folium is used, in terms of raw leaf, that is, when it is prescribed as the weight of raw leaf, in an amount of 0.25 to 400 parts by weight, preferably 0.5 to 200 parts by weight, still preferably 1 to 1 parts by weight, based on 1 part by weight of the extract of *Rosa roxburghii*.

When both *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* are used together with *Rosa roxburghii*, the amounts thereof are not particularly limited. Generally, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. are used in amounts of 0.25 to 400 parts by weight and 0.5 to 800 parts by weight, respectively, in terms of their respective raw leaf, based on 1 part by weight of the extract of *Rosa*

roxburghii. It is preferable that *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. be used in amounts of 0.5 to 200 parts by weight and 1 to 400 parts by weight respectively, still preferably in amounts 1 to 100 parts by weight and 5 to 200 parts by weight respectively.

The pharmaceutical composition or the medicinal agent of the present invention may be administered, with the purpose of the prevention of diseases, as a food having a regulating effect on a living body, i.e., so-called functional food, which can be prepared by adding this composition or agent to food.

When the pharmaceutical composition or the medicinal agent of the present invention comprising one or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* is used as a medicine or so-called health food, it may be in the form of tablet, granule, powder, capsule or syrup. The pharmaceutical composition or the medicinal agent is prepared by mixing the members with a conventional filler or carrier, binder, lubricant or the like and treating the obtained mixture in the conventional manner.

When the pharmaceutical composition or the medicinal agent of the present invention comprising one or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* is used for an animal, such as a mammal, a bird and a fish, especially a livestock, the dosage form thereof is not particularly limited. For example, it may be administered to livestock in a state mixed with a basal diet (or feed or feedstuff). That is, this composition or agent may be mixed with a basal diet (or feed or feedstuff) just before using or may be premixed with a basal diet (or feed or feedstuff). In other words, the feedstuff for an animal comprising a substance selected from the group consisting of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. of the present invention may be administered to an animal as a feed having a biophylactic and regulating function.

The arbitrary basal diet for animals, which is used for preparing the feedstuff according to the present invention, is not particularly limited. Examples of the raw materials constituting the basal diet include grains such as corn, milo and wheat flour, brans such as defatted rice bran and wheat bran, animal substances such as fish meal and skim milk, vegetable oil cake such as soybean oil cake, and additives such as calcium carbonate, calcium phosphate, common salt, vitamin B<sub>2</sub>, DL-methionine, choline chloride, manganese sulfate, dry iron sulfate, calcium iodate, copper sulfate, dry zinc sulfate and sodium saccharin. The basal diet may be prepared by blending some members selected from among these materials. The formulation of the basal diet varies depending upon the animal to which the diet is administered.

The improved feed of the present invention, i.e., the feedstuff containing at least one of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. can be administered to various livestock, poultry, pets and fish. The livestock includes pig, cattle, horse, sheep and rabbit; the poultry include chicken, Japanese quail, turkey and duck; the pets include dog and cat; the fish include yellowtail, sea bream, flatfish, globefish, hardtail, amberjack, salmon, carp, eel, sweetfish, trout and char; and lobsters.

Needless to say, the pharmaceutical composition, the medicinal agent and

the feedstuff of the present invention are non-toxic.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for immunopotentiating and protecting from infectious diseases.

In this case, *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. may be used alone or as a combination of two more of them. In other words, the combination of *Rosa roxburghii* with *Artemisiae argyi folium*, that of *Rosa roxburghii* with *Brassica oleracea* var. *capitata* L., that of *Artemisiae argyi folium* with *Brassica oleracea* var. *capitata* L., and that of *Rosa roxburghii* with *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. can be used.

*Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata* L. exhibit an immunopotentiating and protective activity even when used each alone. However, the use of a combination of two or more of them surprisingly exhibits an immunopotentiating and protective activity which is higher than that of the use of each of them.

The pharmaceutical composition or the medicinal agent of the present invention can safely and effectively potentiate the immune function of livestock such as cattle, pig or horse, a poultry such as chicken or Japanese quail, fish such as young yellowtail, sea bream, eel, trout, or goldfish, a pet such as dog or cat, or a human being to protect the animal or the like from various infectious diseases, which is one of the objects of the present invention.

The term "immunopotentiation" used in this specification means "potentiation of the immune function of an animal such as a human being, mammal, fish or the like".

The pharmaceutical composition or the medicinal agent of the present invention have the effect of enhancing the immune function of an animal including a human being, and serves as a preventive and therapeutic pharmaceutical composition or agent for various maladies and infectious diseases by virtue of this effect, so that the diseases against which the composition or agent is efficacious are not particularly limited. For example, it is efficacious against, e.g., articular rheumatism, autoimmune diseases, bronchial asthma, nutritional disorders, surgical operation, diseases of old age and various infectious diseases such as respiratory infection, sepsis and urinary infection with respect to a human being.

With respect to animals other than humans, this composition or agent is efficacious against scours, epizootic pneumonia, atrophic rhinitis and infectious enterogastritis of a pig, pneumonia and Marek's disease of chicken, scours, pneumonia and mastitis of cattle, and AIDS and leukemia of a pet.

Further, with respect to fish, the infectious disease against which the composition or agent of the present invention is efficacious is not particularly limited and includes bacterial diseases such as streptococcosis and nodosity, and viral diseases.

In this case, the dose of *Rosa roxburghii* to be administered varies depending upon the dosage form and the subject animal, so that it is not particularly limited.

For example, an extract of *Rosa roxburghii* is administered to livestock such as a pig in a dose of 25 mg or above, preferably 50 mg or above, still preferably 100 mg or above per kilogram of the body weight.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for regulating a function of the digestive tract.

In this case, *Rosa roxburghii* is preferably used as the essential component. The activity of regulating the function(s) of the digestive tract, i.e., the gastrointestinal function(s) regulating activity, of pharmaceutical composition or the medicinal agent is synergistically enhanced when the composition or agent further contains *Artemisiae argy folium* or *Artemisiae argyi folium* and *Brassica oleracea* var. *capitata*. The pharmaceutical composition or the medicinal agent is also used as component of a feedstuff, and the feedstuff containing the pharmaceutical composition or the medicinal agent also exhibit the above-described gastrointestinal function regulating activity.

The pharmaceutical composition or the medicinal agent of the present invention can efficaciously and safely regulate the function(s) of the digestive tract of livestock such as cattle, pig or horse, poultry such as chicken or Japanese quail, fish such as young yellowtail, sea bream, trout, carp or goldfish, a pet such as a dog or cat, or humans, which one of the main objects of the present invention.

The term "digestive tract" to be used in this specification refers to organs ranging from the mouth to the anus, particularly to the stomach, duodenum, small intestine, large intestine and rectum.

The term "function(s) of the digestive tract" or "gastrointestinal function(s)" used in this specification means digestion, absorption and digestive motion such as peristalsis. The pharmaceutical composition or the medicinal agent for the digestive tract according to the present invention is useful as a therapeutic and preventive pharmaceutical composition or agent for diarrhea when it acts on the lower part of the digestive tract, such as the large intestine or rectum, while it is efficacious in alleviating a sharp pain and in moderating gastric emptying and borborygmus when it acts on the upper part thereof, such as the stomach or duodenum.

When the pharmaceutical composition or the medicinal agent for regulating the function(s) of the digestive tract is used as a therapeutic and preventive pharmaceutical composition or agent for diarrhea, it is efficacious against various diarrheas which are not limited in cause. Examples of the diarrhea include bacterial diarrhea such as salmonella, viral diarrhea such as caused by adenovirus, parasitic diarrhea such as amebic dysentery, toxic diarrhea such as food and drug poisoning, allergic diarrhea caused by, e.g., food allergy, functional diarrheas such as constipation and neurotic diarrhea, diarrheas caused by the use of an antibiotic (such as one caused by microbial substitution and staphylococcal diarrhea), and chronic diarrheas caused by the disorder of digestion and absorption, excess gut hormone and colic diseases.

With respect to chickens and pigs, the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is efficacious

against bacterial diarrhea such as caused by *Escherichia coli* or swine dysentery, viral diarrhea caused by TGE or rotavirus, or simple diarrhea such as one caused by stress or dietetic diarrhea. Further, it is efficacious also against various diarrheas of fish.

In this case, the dose of *Rosa roxburghii* to be administered varies depending upon the dosage form and the subject animal, so that it is particularly limited.

For example, an extract of *Rosa roxburghii* is administered to livestock such as a pig in a dose of 20 mg or above, preferably 40 mg or above, still preferably 100 mg or above per kilogram of the body weight.

It will be described about the case that the pharmaceutical composition the medicinal agent or the feedstuff of the present invention is used improving antibiotic absorption.

In this case, the pharmaceutical composition or the medicinal agent comprising *Rosa roxburghii* or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata L.* is administered simultaneously with an antibiotic or before or after the administration of an antibiotic. As pharmaceutical composition or the medicinal agent, *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata L.* may be administered by itself in a raw, dried or pulverized state, or a formulation may be administered. Alternatively, the pharmaceutical composition or the medicinal agent may be added to food or feed in a conventional manner. Namely, the feedstuff comprising a basal diet and least one of *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata L.* of the present invention may be used.

The feedstuff of the present invention is prepared by adding *Rosa roxburghii* or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata L.* to the basal diet described above. The amount of *Rosa roxburghii*, or the combination, is preferably 0.001% by weight or above still preferably 0.01% by weight or above in terms of the dried extract thereof, i.e., as the weight of the dried extract thereof.

The pharmaceutical composition, the medicinal agent or the feedstuff for improving antibiotic absorption according to the present invention may be administered simultaneously with an antibiotic or, before or after the administration of an antibiotic. In any case, the effect of improving absorption of an antibiotic can be attained.

In this case, the dose of *Rosa roxburghii* or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata L.* is not particularly limited but varies depending upon the dosage form, the subject animal or the dose of the antibiotic.

For example, when the pharmaceutical composition or the medicinal agent for improving antibiotic absorption comprising two or more members selected from among *Rosa roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata L.* is administered to livestock such as a pig a powdered extract, the powdered extract is administered in an amount of part by weight or above, preferably 1 part by weight or above, still



preferably 3 parts by weight or above based on 1 part by weight of the antibiotic administered. When *Rosa roxburghii* is administered alone, the amount thereof may be the same as that described above.

It will next be described when the pharmaceutical composition, the medicinal agent or the feedstuff of the present invention is used for accelerating growth of an animal or improving an egg production rate, weight, egg quality or eggshell strength.

One of the objects of the present invention is to provide a novel pharmaceutical composition, medicinal agent or feedstuff for accelerating growth of an animal, a novel pharmaceutical composition, medicinal agent or feedstuff for improving the egg production rate, egg weight, egg quality or eggshell strength of animals, and a method for administering them to animals.

More specifically, one of the objects of the present invention is to provide a novel pharmaceutical composition, medicinal agent or feedstuff which can accelerate an increase in the body weight and can improve the survival rate, feed conversion ratio of animals, and a novel pharmaceutical composition, medicinal agent or feedstuff which can improve the survival rate, egg production rate, egg weight, egg quality, eggshell strength and feed conversion ratio of animals.

The term "growth acceleration" or "accelerating growth of an animal" as used herein includes also growth improvement and growth acceleration of fetuses. The expression "improvement in the egg production rate, egg weight, egg quality and eggshell strength" means an improvement in the production rate, an increase in the egg weight, and improvements in the eggshell and egg quality.

In this case, *Rosa roxburghii* alone or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisia argyi* folium and *Brassica oleracea* var. capitata L., preferably a combination of three members described above, as the active ingredient is administered to animals. *Rosa roxburghii*, *Artemisia argyi* folium and *Brassica oleracea* var. capitata L. can be administered directly to animals or they can be incorporated in the feedstuff or feed by an ordinary method. Namely, the feedstuff comprising a basal diet and at least one of *Rosa roxburghii*, *Artemisia argyi* folium and *Brassica oleracea* var. capitata L. of the present invention may be used.

The feedstuff of the present invention is prepared by adding *Rosa roxburghii* alone or a combination of two or more members selected from among *Rosa roxburghii*, *Artemisia argyi* folium and *Brassica oleracea* var. capitata L. to the basal diet described above. The amount of *Rosa roxburghii* or the combination is preferably 0.001% by weight or above, still preferably 0.01% by weight or above in terms of the dried extract thereof, i.e., as the weight of the dried extract thereof.

The pharmaceutical composition, the medicinal agent or the feedstuff accelerates the growth of animals, and improves the raising or feed conversion ratio. When it is administered to birds and fish, it results in an improvement in the egg production rate, egg weight, egg quality or eggshell strength of them.

The dose of *Rosa roxburghii* is not particularly limited in this case,

since it varies depending on the dosage form thereof and the subject animal.

When *Rosa roxburghii* extract is administered to livestock, such as cat and pigs, the amount thereof is usually at least 100 mg, preferably at least 50 mg and still preferably at least 100 mg, based on 1 kilogram the body weight thereof.

#### DETAILED DESCRIPTION:

#### EXAMPLES

The present invention will now be described in more detail with reference to the following Examples which should not be considered to limit the scope of the present invention.

In the Examples, the description of the dose of each component, e.g., mg/kg p.o." means oral administration in a dose of 10 mg per kilogram body weight. Further, the symbols "\*", and "\*\*\*" in the tables mean  $p < 0.05$  and  $p < 0.01$ , respectively, in the X.sup.2 test.

The *Rosa roxburghii* used in Examples A-1, B-1 and B-2 is one commercially available under the trade name of "Rosa roxburghii extract powder MF", which is a product of Maruzen Seiyaku K. K. comprising 30% of an extract of *Rosa roxburghii* and 70% dextrin. The dose of *Rosa roxburghii* is expressed in terms of the weight of this extract powder. The dose of *Artemisiae argyi* folium is expressed in terms of the weight of an extract thereof having 4-fold concentration with respect to its normal weight, while the dose of *Brassica oleracea* var. capitata L. is expressed in terms of the weight of an extract thereof having a 9-fold concentration with respect to its normal weight.

#### EXAMPLE A-1

As shown in Tables A-1 and A-2, *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. capitata L. were each orally administered (Table A-1), or as a mixture of two or more of them (Table A-2), to the SLC:ICR male mice (age: 5 to 6 weeks, weight: 25 to 33 g) in doses of 100 to 2000 mg/kg, while physiological saline was orally administered thereas a control. After 24 hours, clinically available *Escherichia coli* (5.0.times.10.sup.7 CFU/mouse, 0.2 ml) was intravenously inoculated in each mouse to determine the survival rate based on the number of viable mice after 7 days from the infection. The results are given in Tables A-1 and A-2.

TABLE A-1	Survival	Sample rate
x.sup.2 test	control (physiological saline p.o.)	
0 <i>Artemisiae argyi</i> folium 500 mg/Kg p.o.	20	<i>Artemisiae argyi</i> folium 1000 mg/Kg p.o. 30
<i>Artemisiae argyi</i> folium 1000 mg/Kg p.o.	30	<i>Artemisiae argyi</i> folium 2000 mg/Kg p.o. 50 *
<i>Brassica oleracea</i> var. capitata L. 10	500 mg/Kg p.o.	<i>Brassica oleracea</i> var. capitata L. 30
1000 mg/Kg p.o.	<i>Brassica oleracea</i> var. capitata L. *	2000 mg/Kg p.o.
<i>Rosa roxburghii</i> 500 mg/Kg p.o.	20	<i>Rosa roxburghii</i> 1000 mg/Kg p.o. 30
<i>Rosa roxburghii</i> 2000 mg/Kg p.o.	50	*

#### TABLE A-2

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Sample Survival rate (%) x.sup.2 test

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control (physiological saline p.o.) 0 Artemisiae argyi folium 500 mg/Kg p.o. and 40 \* Brassica oleracea var. capitata L. 500 mg/Kg p.o. Artemi argyi folium 1000 mg/Kg p.o. and 70 \* Brassica oleracea var. capitata 1000 mg/Kg p.o. Artemisiae argyi folium 2000 mg/Kg p.o. and 100 \*\* Brassica oleracea var. capitata L. 2000 mg/Kg p.o. Rosa roxburghii 500 mg/Kg p.o. and 60 \* Artemisiae argyi folium 500 mg/Kg p.o. Rosa roxbur 1000 mg/Kg p.o. and 90 \*\* Artemisiae argyi folium 1000 mg/Kg p.o. Rosa roxburghii 2000 mg/Kg p.o. and 100 \*\* Artemisiae argyi folium 2000 mg/p.o. Rosa roxburghii 500 mg/Kg p.o., 70 \* Artemisiae argyi folium 500 mg/Kg p.o. and Brassica oleracea var. capitata L. 500 mg/Kg p.o. Rosa roxburghii 1000 mg/Kg p.o., 100 \*\* Artemisiae argyi folium 1000 mg/Kg and Brassica oleracea var. capitata L. 1000 mg/Kg p.o.

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As shown in Table A-1, the survival rate increases depending upon the of Rosa roxburghii, Artemisiae argyi folium or Brassica oleracea var. capitata L., which reveals that these plants have a protective effect.

As shown in Table A-2, the simultaneous use of Rosa roxburghii with Artemisiae argyi folium, that of Artemisiae argyi folium with Brassica oleracea var. capitata L. and that of Rosa roxburghii with Artemisiae argyi folium and Brassica oleracea var. capitata L. exhibit the protec effect exceeding the arithmetic sum of the respective effects of these plants. That is, the above simultaneous use exhibits a significant synergistic effect, being of great value.

The above test in Example A-1 reveals that the medicinal agent for immunopotentiating and protecting from infectious diseases of the pres invention has an excellent protective activity against Gram-negative bacteria, which suggests that the agent of the present invention is no only an agent which exhibits a low antimicrobial activity only against Gram-positive bacteria, like Artemisiae argyi folium, but is also an a which potentiates the immune function itself. Accordingly, the agent o the present invention is effective in the prevention and treatment of various maladies and useful as a preventive and therapeutic agent for various infectious diseases or as a functional food or feed. Thus, the present invention is of great value.

#### EXAMPLE B-1

As shown in Table B-1, Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were each orally administered to si SLC:SD male rats (age: 6 weeks, weight: 250 to 300 g) in a dose of 125 500 mg/kg, while physiological saline was orally administered thereto control. After 30 minutes, 1 ml of castor oil was further orally administered to the rats to determine the time which elapsed until the occurrence of diarrhea and the total amount of excrement given within hours of the administration. The antidiarrheal index of each case was calculated from these values according to the following formula 1 to evaluate the antidiarrheal activity. The measured values and the antidiarrheal index are given in Table B-1. ##EQU1##

TABLE B-1	<hr/>	Time until the Total
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occurrence amount of Anti- of diarrhea excrement	diarrheal Sample (min)	control (physiologica
(g) index		
38.5 .+-. 5.9 6.7 .+-. 0.67 1.00 saline p.o.)	Artemisiae argyi	53.3 .+
13.8 4.5 .+-. 0.83 2.06 folium 125 mg/Kg p.o.	Artemisiae argyi	71.8 .+
18.4 3.7 .+-. 1.09 3.38 folium 250 mg/Kg p.o.	Artemisiae argyi	95.8 .+
12.8 2.0 .+-. 0.68 8.33 folium 500 mg/Kg p.o.	Brassica oleracea var.	4
.+-. 7.9 5.9 .+-. 1.16 1.36 capitata L. 125 mg/Kg p.o.	Brassica olerac	
var. 49.5 .+-. 10.7 4.7 .+-. 1.14 1.83 capitata L. 250 mg/Kg p.o.	Bras	
oleracea var. 51.2 .+-. 8.2 3.8 .+-. 1.21 2.34 capitata L. 500 mg/Kg p		
Rosa roxburghii 50.5 .+-. 10.0 4.2 .+-. 1.67 2.09 125 mg/Kg p.o.	Rosa	
roxburghii 68.3 .+-. 6.4 3.7 .+-. 0.80 3.21 250 mg/Kg p.o.	Rosa roxbur	
77.8 .+-. 14.4 1.8 .+-. 0.70 7.52 500 mg/Kg p.o.		

It was ascertained from the results given in Table B-1 that Artemisiae argyi folium, Rosa roxburghii and Brassica oleracea var. capitata L. h an antidiarrheal activity.

#### EXAMPLE B-2

As shown in Tables B-2 and B-3, two or more members selected from amon Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were orally administered to six SLC:SD male rats (age: 6 weeks, weight: 250 to 300 g) each in doses of 125 to 500 mg/kg, while physiological saline was orally administered thereto as a control. Aft 30 minutes, 1 ml of castor oil was further administered orally to the to determine the time elapsed until the occurrence of diarrhea and the total amount of the excrement given within 2 hours of the administrati The antidiarrheal index of each case was calculated from these values according to the above formula 1 to evaluate the antidiarrheal activit The measured values and the antidiarrheal index are given in Tables B- and B-3.

TABLE B-2	Time until the Total
occurrence amount of Anti- of diarrhea excrement	diarrheal Sample (min)
(g) index	control (physiologica
38.5 .+-. 5.9 6.7 .+-. 0.67 1.00 saline p.o.)	Artemisiae argyi 62.7 .+
13.4 4.2 .+-. 1.02 2.60 folium 125 mg/Kg p.o. and Brassica oleracea va	capitata L. 125 mg/Kg p.o. Artemisiae argyi 83.5 .+-. 15.3 3.2 .+-. 1.
4.54 folium 250 mg/Kg p.o. and Brassica oleracea var. capitata L. 250	mg/Kg p.o. Artemisiae argyi 99.3 .+-. 13.2 1.8 .+-. 0.70 9.59 folium 5
mg/Kg p.o. and Brassica oleracea var. capitata L. 500 mg/Kg p.o. Rosa	roxburghii 78.4 .+-. 8.6 3.5 .+-. 1.73 3.89 125 mg/Kg p.o. and Artemis
argyi folium 125 mg/Kg p.o. Rosa roxburghii 90.3 .+-. 12.4 2.6 .+-. 1.	6.04 250 mg/Kg p.o. Artemisiae argyi folium 250 mg/Kg p.o. Rosa roxbur
101.8 .+-. 14.3 1.6 .+-. 1.58 11.06 500 mg/Kg p.o. Artemisiae argyi fo	500 mg/Kg p.o.

TABLE B-3	Time until the Total
occurrence amount of Anti- of diarrhea excrement	diarrheal Sample (min)
(g) index	control (physiologica
38.5 .+-. 5.9 6.7 .+-. 0.67 1.00 saline p.o.)	Rosa roxburghii 91.5 .+
17.4 3.5 .+-. 1.73 6.37 125 mg/Kg p.o. Artemisiae argyi folium 125 mg/	p.o. and Brassica oleracea var. capitata L. 125 mg/Kg p.o. Rosa roxbur
102.0 .+-. 13.0 1.8 .+-. 0.70 9.85 250 mg/Kg p.o. Artemisiae argyi fol	250 mg/Kg p.o. and Brassica oleracea var. capitata L. 250 mg/Kg p.o. R
roxburghii 110.8 .+-. 14.0 1.3 .+-. 1.16 14.82 500 mg/Kg p.o. Artemisi	

argyi folium 500 mg/Kg p.o. and Brassica oleracea var. capitata L. 500 mg/Kg p.o. \_\_\_\_\_

It has been ascertained from the results given in Tables B-1 and B-2 that the simultaneous use of Rosa roxburghii with Artemisiae argyi folium exhibits a synergistic antidiarrheal activity, while it has been ascertained from the results given in Tables B-1 and B-3 that the simultaneous use of Rosa roxburghii with Artemisiae argyi folium and Brassica oleracea var. capitata L. also exhibits a synergistic antidiarrheal activity.

The above test in Examples B-1 and B-2 reveals that the medicinal agent for regulating a function of a digestive tract according to the present invention has an excellent antidiarrheal activity. Accordingly, the medicinal agent for regulating a function of a digestive tract of the present invention is effective in the prevention and treatment of diarrhea caused by various diseases, and is useful as a preventive or therapeutic agent for diarrhea or a functional food or feed. Thus, the present invention is of great value.

The mixture of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. used in the following Examples C-1, C-2, D-1 D-6 were prepared as follows:

Raw leaves (3 kg) of Artemisiae argyi folium were immersed in 15 l of water and the obtained mixture was boiled for 30 minutes to conduct extraction. The resulting mixture was filtered to recover a filtrate. The filtrate was concentrated to 1.8 kg by heating to give an extract. This extract was powdered by spray drying to give a dry powder of the extract of Artemisiae argyi folium. Similarly, Brassica oleracea var. capitata L. was also converted into a dry powder of the extract thereof. 1500 g of Rosa roxburghii extract powder MF (a product of Maruzen Seiyaku K. K. comprising 30% of an extract of Rosa roxburghii and 70% of dextrin) was mixed with 675 g of the above dry powder of the extract of Artemisiae argyi folium and 225 g of the above dry powder of the extract of Brassica oleracea var. capitata L. to give 2400 g of a mixed powder (a weight ratio of Rosa roxburghii:Artemisiae argyi folium:Brassica oleracea var. capitata L. being 2:3:1).

#### EXAMPLE C-1

The above mixture of the dry powders or each of the dry powders was orally administered to four SLC:ICR male mice (age: 5 to 6 weeks, weight: 25 to 33 g) in a dose given in Table C-1 together with 10 mg per kilogram of body weight of amoxicillin (hereinafter referred to simply as "AMPC") an antibiotic. After 2 hours, blood was gathered from the mice to determine the serum AMPC concentrations, which were compared with those of mice to which AMPC alone had been administered. Each serum AMPC concentration was determined by using a strain of *Sarcina lutea* (Gram-positive and AMPC-sensitive bacterium) according to the disk method (this method of measurement was also employed in Example C-2).

The results are given in Table C-1.

TABLE C-1	Serum AMPC concn.
.µg/ml (figures in parentheses being ratio thereof to the concn. of control)	AMPC 10 mg/kg (control)

2.12  $\pm$  1.14 (1.00) AMPC 10 mg/kg + a mixed 2.70  $\pm$  1.22 (1.27) po of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. 10 mg/kg AMPC 10 mg/kg + a mixed 3.04  $\pm$  1.22 (1.43) pow of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. 30 mg/kg AMPC 10 mg/kg + a mixed 3.11  $\pm$  1.97 (1.47) pow of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. 90 mg/kg AMPC 10 mg/kg + a mixed 3.53  $\pm$  1.93 (1.67)\* po of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. 190 mg/kg AMPC 10 mg/kg + a mixed 5.62  $\pm$  0.57 (2.65)\*\* powder of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. 2000 mg/kg AMPC 10 mg/kg + Rosa 4.87  $\pm$  1.81 (2.30) roxburghii 10 mg/kg AMPC 10 mg/kg + Artemisiae 2.95  $\pm$  1.13 (1.39) a folium 10 mg/kg AMPC 10 mg/kg + Brassica 2.20  $\pm$  1.11 (1.04) oleracea var. capitata L. 10 mg/kg

As shown in Table C-1, the group of the mice to which AMPC and Rosa roxburghii had been administered and the groups of the mice to which A Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. had been administered were apparently improved in the absorption of AMPC depending upon the dose thereof as compared with the group of the mice to which AMPC alone had been administered.

#### EXAMPLE C-2

AMPC (10 mg/kg) alone or the mixture of AMPC (10 mg/kg) with the mixed powder (30 mg/kg) comprising Rosa roxburghii, Artemisiae argyi folium Brassica oleracea var. capitata L. was administered to six SPF male pig (age: 1 month, weight: about 15 kg) mixed with a basal diet to gather blood from the pigs after 0.5, 1, 2, 3 and 6 hours. The serum AMPC concentrations of the pigs to which AMPC and the mixed dry powder had administered were determined and compared with those of the pigs to which AMPC alone had been administered.

The results are given in Table C-2.

The formulation of the above basal diet are given in Table C-3.

TABLE C-2

after after after after after 0.5 hr 1 hr 2 hrs 3 hrs 6 hrs																
AMPC 10 mg/kg (control)		3.07	+-	1.55	6.03	+-	0.80	3.71	+-	0.63	2					
		+-	0.40	0.85	+-	0.75	AMPC 10 mg/kg + a mixed powder		6.03	+-	1.49					
		7.81	+-	1.49	5.85	+-	0.80	3.71	+-	1.41	1.74	+-	1.16	of Rosa		
roxburghii, Artemisiae argyi		(1.63)**			(1.30)*			(1.58)**		(1.77)*	(2.05)*					
folium and Brassica oleracea var. capitata L.		30 mg/kg														

Note) Serum AMPC concentration,  $\mu$ g/ml The figures in parentheses are the ratios of the serum AMPC concentration to that of the control.

TABLE C-3	Initiation of															
Administration feeding to 5	5	to 7 weeks	period	weeks	of age	of age										
Component	Amt. (%)	Amt. (%)														par
wheat	33.8	20.0	glucose	9.0	5.0	sugar	3.0	--	wheat flour	--	20.0	yellow				
corn	--	23.6	skimmilk powder	40.0	10.0	fish meal	3.0	5.0	soybean oil	c						
	3.0	9.0	Torula yeast	2.0	2.0	soybean oil	4.0	3.0	common salt	0.2	0.2					
tricalcium phosphate	1.2	1.4	mineral mixture	0.2	0.2	vitamin mixture	0									

0.2 flavor, antibiotic and 0.4 0.4 other feed additives

As shown in Table C-2, the pigs to which AMPC and the mixed powder had been administered were apparently improved in the absorption of AMPC o a period of 0.5 to 6 hours after the administration as compared with t pigs to which AMPC alone had been administered.

The results of the above tests suggest that the medicinal agent for improving antibiotic absorption and the feedstuff according to the pre invention are so efficacious in improving the absorption of an antibio that the dose of the antibiotic can be reduced. Accordingly, the prese invention is of great value.

#### EXAMPLE D-1

40 sound pigs (castrated pigs of about 10 days old; 3.0 to 3.5 kg) wer divided into four groups, i.e., three groups each consisting of ten pi to which the mixed powder comprising *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and a control group consisting of ten pigs to which an ordinary basal diet f of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered. A feedstuff enriched with the mixed powd of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. in amounts of 0.03, 0.1 or 0.3% by weight was administered the pigs of one of the three groups for 30 days. After 30 days, the in of the feedstuff and the gain of body weight were determined to calcul the rate of raising and the feed conversion ratio (feedstuff intake/ga of body weight). The results were compared with those of the control group.

The composition of the artificial milk-based basal diet for piglings w as shown in Table D-1.

The results of the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and th control group were compared with each other. The t-test was conducted the gain of body weight in the control group and the groups of the pre invention. Table D-2 shows the results.

TABLE D-1										Initiation of feeding	
5 5 to 7 weeks Feeding period weeks of age of age										Component Amt. (%)	
Amt. (%)										Roast wheat 33.8 20.0	
Glucose	9.0	5.0	Sugar	3.0	--	Wheat flour	--	20.0	Yellow corn	--	23.6
milk powder	40.0	10.0	Fish meal	3.0	5.0	Soybean oil	cake	3.0	9.0	Torul	
yeast	2.0	2.0	Soybean oil	4.0	3.0	Common salt	0.2	0.2	Tricalcium phosp		
1.2	1.4	Mineral mixture	0.2	0.2	Vitamin mixture	0.2	0.2	Flavor, antibi			
and 0.4	0.4	other feed additives									

TABLE D-2										Gain of Feedstuff bod	
Feed intake weight conversion (A) (B) ratio Rate of Group (kg/pig)											
(kg/pig) (A/B) raising										Test gro	
11.24	5.3	+-.	1.33	2.12	88.8%	(0.03%)	** Test group	11.79	6.3	+-.	1.
1.87	93.0%	(0.1%)	** Test group	12.32	5.7	+-.	1.04	2.16	100%	(0.3%)	*
Control	8.50	3.4	+-.	1.15	2.50	81.9%	group				

The gain of body weight per pig in the groups to which *Rosa roxburghii* *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered was larger than that in the control group, and the feed conversion ratio and the rate of raising were increased in the former group.

#### EXAMPLE D-2

18 bull Holstein calves of about 7 days old were divided into three groups, i.e., two groups each consisting of six calves to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and a control group consisting of six calves to which neither of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was administered. The mixed powder of *Rosa roxburghii* *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. was mixed into a milk substitute for raising suckling calves and into a feedstuff containing a milk substitute (artificial milk) for raising suckling calves in an amount of 0.125 or 0.25% by weight based on the basal diet. The milk substitute or the feedstuff was repeatedly administered for 20 days. The dose of the mixed powder was 2.5 g/day or 5 g/day per calf. Each calf was weighed at the beginning of the test and after the completion of the test (after 20 days) to calculate the gain of body weight and the rate of body weight gain per day.

The composition of the feedstuff containing the milk substitute for raising suckling calves was as shown in Table D-3.

The results of the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and those of the control group are compared with each other. The t-test was conducted in the control group and the groups of the present invention.

Table D-4 shows the results.

TABLE D-3				Component Amt. (%)	
				Skim milk powder	60.0
Animal fat or oil	20.0	Fish soluble	4.0	Dry whey	
mixture	0.7	Vitamin mixture	0.5	Mineral	

TABLE D-4		Gain of body weight		Gain of body weight	
Group	(kg/calf)	Group	(kg/calf/day)	Group	(kg/calf/day)
Test group	41.0	Test group	42.5	Control group	25.7
2.5 g/day	0.29	5 g/day	0.59	Control group	0.36
2.5 g/day	0.29	5 g/day	0.59	Control group	0.36

The gain of body weight per day in the groups to which *Rosa roxburghii* *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered was larger than that in the control group, and the rate of body weight gain was increased in the former group.

#### EXAMPLE D-3

80 pullet broilers (chunky) were divided into two groups, i.e., a group consisting of 40 broilers to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and a control group consisting of 40 broilers to which neither of *Rosa*



roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata was administered.

The mixed powder of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. was administered in an amount of 0.1% by weight based on the basal diet for broilers to the group, to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata were administered, for 6 weeks. Each broiler was weighed at the beginning of the test and after 6 weeks. The average gain of body weight in the group to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were administered and that in the control group were calculated.

The composition of the basal diet for broilers was as shown in Table D

The results of the group to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were administered and those of the control group are compared with each other. The x<sup>2</sup>-test for the average gain of body weight was conducted in the control group and the group to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were administered.

Table D-6 shows the results.

Initiation of feeding									
to 5 weeks		to 6 weeks		to 3 weeks		weeks of Feeding		period of age	
Component	Amt. (%)	Amt. (%)	Amt. (%)	Amt. (%)	Amt. (%)	Amt. (%)	Amt. (%)	Amt. (%)	Amt. (%)
Corn	50.45	55.30	55.34	Milo	10.15	15.01	Soybean oil cake	18.0	13.0
Rapeseed oil cake	--	2.5	2	Fish meal	8.0	5.0	5.01	Fish soluble	2.0
Absorbent feed	3.0	--	--	Gluten meal	2.0	--	--	Torula yeast	1.0
Meat/bone meal	--	3.0	3.0	Raw rice bran	--	2.0	2.0	Animal fat or oil	3.3
2.8	2.81	Com salt	0.25	0.25	0.25	Calcium carbonate	0.6	0.3	0.30
Dicalcium phosphate	0.5	0.5	Lysine	--	--	--	Methionine	0.18	--
Vitamin mixture	0.1	0.1	Choline chloride	0.05	0.06	0.06	Mineral mixture	0.1	0.1
Flavor, antibiotic and	0.17	0.09	--	other feed additives					

Av. body wt. at initiation				Av. body wt. Av. gain of of feeding after 6 weeks			
Group (g/broiler)	(g/broiler)	(g/broiler)	(g/broiler)	Test group	43.8	+-	0.08
2065	8.7	2021.7	+-	8.6**	(0.1%)	Control group	43.6
1879	1835.4	+-	10.1				

The increase in the average body weight in the group to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata were administered was larger than that in the control group.

#### EXAMPLE D-4

12,000 young yellowtails having an average body weight of about 600 g divided into two groups, i.e., a group consisting of 6,000 young yellowtails to which Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. were administered and a control group consisting of 6,000 young yellowtails to which neither of Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. was administered.

mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were mixed with a basal diet for young yellowtails (moist pellets) in an amount of 0.5% by weight based on the basal diet and the resultant mixture was administered to them at intervals of three days for one month. The results were compared with those of the control group. The young yellowtails were raised in a crawl having a surface of about 3 m.times.3 m.times.3 m. During the test period, the feedstuff was administered once a day in an amount of about 3% by weight based on the body weight a day.

The composition of the basal diet for young yellowtails was as shown in Table D-7.

The death rate during the test period in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* were administered and the death rate during the test period in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered and that in the control group were compared with each other to evaluate the effect of improving the rate of raising. The x.sup.2 -test was conducted in the control group and the group of the present invention. The results are given in Table D-8.

TABLE D-7		Component Amt. (%)	
		Fish meal	56.0
		Meat/bone meal	3
Soybean oil cake	5.0	Corn gluten meal	3.0
Torula yeast	2.0	Wheat flour	28.4
Vitamin mixture	1.0	Choline chloride	0.3
Inorg. mixture	1.0	Sodium polyacrylate	0.3

TABLE D-8		No. of No. of died	
samples	samples	Group (fish)	(fish)
Test group (0.1%)	6000	180	**
Control group	6000	750	

The death rate during the test period in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* were administered was far smaller than that in the control group to prove the effect of improving the rate of raising.

*Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. used in pellet for mice and rats, of a preparation prepared by spraying an aqueous suspension of the mixed extract powder, followed by drying, was used to form the pellet for mice and rats. Each feedstuff containing *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* thus prepared was administered to each group of male slc:SD rats, which consisted of 8 rats of 21 days old (body weight: 50 to 57 g), for one month and then each rat was weighed to calculate the rate of body weight gain.

The composition of the basal diet for rats (pellet) was as shown in Table D-9.

The body weights of the rats weighed during the test period in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. were administered were compared with those in the control group to examine the effect on the gain of body weight in both groups. t-test was conducted in the control group and the groups to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata*

L..

The results are given in Table D-10.

TABLE D-9				Component Amt. (%)			
				Casein	14.12	Corn starch	58.40
Glucose	12.83	Soybean oil	5.45	Cellulose	5.00	Minerals	4.00
Vitamins A							
D	0.10	Vitamin B	0.10				

TABLE D-10

Average body weight (g) and rate of body weight gain in each group Gro  
week 1 week 2 weeks 3 weeks 4 weeks

Test group	52.9	+-.	1.48	106.7	+-.	4.29	162.4	+-.	7.17	213.4	+-.	6
	258.2	+-.	7.70	(0.03%)	(1.00)	(2.02)	(3.06)	(4.03)	(4.86)	*	Test grou	
	53.2	+-.	1.83	107.6	+-.	3.77	162.6	+-.	7.28	216.3	+-.	9.41
	10.96	(0.1%)	(1.00)	(2.02)	(3.06)	(4.07)	(4.88)	*	Test group	53.7	+-.	
	2.65	107.6	+-.	3.23	166.7	+-.	6.09	221.4	+-.	10.61	270.2	+-.
	(0.3%)	(1.00)	(2.00)	(3.10)	(4.12)	(5.03)	**	Control group	53.8	+-.	1	
	102.9	+-.	5.23	157.3	+-.	9.91	204.5	+-.	13.41	241.9	+-.	17.08
	(1.91)	(2.92)	(3.80)	(4.50)								

After the administration of *Rosa roxburghii*, *Artemisiae argyi* folium a *Brassica oleracea* var. *capitata* L., an increment in the rate of body weight gain of 0.23 to 0.53 was recognized depending on the dose, as compared with the control group.

## EXAMPLE D-6

The tests were conducted twice by using groups of Dekalb-TX chickens o 182 days old (average body weight: 1.5 kg). Each group consisted of 8 chickens. The average results are given in Table D-11 and D-12 . In th test, the chickens were divided into three groups, i.e., a group to wh 0.03% of the mixed powder of *Rosa roxburghii*, *Artemisiae argyi* folium *Brassica oleracea* var. *capitata* L. was administered, a group to which of the mixed powder was administered, and the control group. The feedi was conducted for a period of 18 days.

After the completion of the feeding, body weight and egg weight determination, egg quality test and eggshell strength determination (w an eggshell strength meter) were conducted. Further, the feed conversi ratio (feedstuff intake/gain of body weight) was calculated.

The results are given in Tables D-11 and D-12.

TABLE D-11				Egg Feed prodn. Egg			
weight/day	conversion	Group rate	(g/chicken.day)	ratio			
				Test group	79.7%	47.3	2.14
Test group	79.3%	46.6	2.28	(0.1%)	Control	74.2%	44.2
				Note) Egg weight/day indicates			
				total weight of eggs laid by a chicken in a day.			

TABLE D-12			Eggshell		Eggshell		
strength	thickness	Eggshell	Group	(kg/cm.sup.2)	(mm)	(wt.%)	

									Test group	3.40	0.358	10.6	(0.0)
Test group	3.16	0.359	10.4	(0.1%)	Control	3.12	0.374	10.5	group				

It will be apparent from Table D-11 that the egg production rate, egg weight and feed conversion ratio were increased more remarkably in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. had been administered than those in the cont group.

It will be apparent from Table D-12 that as compared with the thickness of the eggshell, the eggshell strength was more increased in the group to which *Rosa roxburghii*, *Artemisiae argyi* folium and *Brassica oleracea* var. *capitata* L. had been administered than that in the control group. The results promise the prevention of egg cracking.

One embodiment of the present invention provides a medicinal agent or animal feedstuff for accelerating growth of an animal which comprises roxburghii alone or Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. to be administered to animals directly or as additives to be incorporated into a basal diet so as to improve the body weight gain, survival rate and feed conversion ratio of an animal; and another embodiment of the present invention provides a medicinal agent improving an egg production rate, egg weight, egg quality or eggshell strength of animals. Since Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata L. are used, a high degree of safety is afforded and no environmental pollution is caused. Further, Rosa roxburghii, Artemisiae argyi folium and Brassica oleracea var. capitata are available at low costs and are excellent as a medicinal agent or animal feedstuff for accelerating growth of an animal and a medicinal agent for improving an egg production rate, egg weight, egg quality or eggshell strength of an animal and have a high storageability.

The invention being thus described, it will be obvious that the same may be varied in many ways. Such variations are not to be regarded as a departure from the spirit and scope of the invention, and all such modifications as would be obvious to one skilled in the art are intended to be included within the scope of the following claims.

CLAIMS:

What we claim is:

1. A method for increasing the egg production rate, egg weight or eggs strength of a bird, which comprises administering a pharmacologically effective amount of a composition comprising a mixture of the plants *Roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* to said bird.
2. The method according to claim 1, wherein the bird is a chicken.
3. The method according to claim 1, wherein the composition is raw, in dried state or an extract.
4. The method according to claim 1, wherein the composition is in a dry state.

5. The method according to claim 1, wherein the composition is an extr

6. A method for increasing the egg production rate, egg weight or eggs strength of a fish, which comprises administering a pharmacologically effective amount of a composition comprising a mixture of the plants *Roxburghii*, *Artemisiae argyi folium* and *Brassica oleracea var. capitata* to said fish.